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(57) Abstract

The present invention relates to an isolated protein or polypeptide corresponding to a protein or polypeptide of a Rupestris stem pitting associated virus. The encoding DNA molecule, either alone in isolated form, in an expression system, a host cell, or a transgenic grape plant, is also disclosed. Other aspects of the present invention relate to a method of imparting Rupestris stem pitting associated virus resistance to grape plants by transforming them with the DNA molecule of the present invention, and a method of detecting the presence of a Rupestris stem pitting associated virus, such as RSPaV-1, in a sample.

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RUPESTRIS STEM PITTING PITTING ASSOCIATED VIRUS NUCLEIC ACIDS, PROTEINS, AND THEIR USES

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FIELD OF THE INVENTION

The present invention relates to *Rupestris* stem pitting associated virus ("RSPaV") proteins, DNA molecules encoding these proteins, and diagnostic and other uses thereof.

BACKGROUND OF THE INVENTION

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The world's most widely grown fruit crop, the grape (Vitis sp.), is cultivated on all continents except Antarctica. However, major grape production centers are in European countries (including Italy, Spain, and France), which constitute about 70% of the world grape production (Mullins et al., Biology of the Grapevine, Cambridge, U.K.:University Press (1992)). The United States, with 300,000 hectares of grapevines, is the eighth largest grape grower in the world. Although grapes have many uses, a major portion of grape production (~80%) is used for wine production. Unlike cereal crops, most of the world's vineyards are planted with traditional grapevine cultivars, which have been perpetuated for centuries by vegetative propagation. Several important grapevine virus and virus-like diseases, such as grapevine leafroll, corky bark, and Rupestris stem pitting ("RSP"), are transmitted and spread through the use of infected vegetatively propagated materials. Thus, propagation of certified, virus-free materials is one of the most important disease control measures. Traditional breeding for disease resistance is difficult due

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to the highly heterozygous nature and outcrossing behavior of grapevines, and due to polygenic patterns of inheritance. Moreover, introduction of a new cultivar may be prohibited by custom or law. Recent biotechnology developments have made possible the introduction of special traits, such as disease resistance, into an established cultivar without altering its horticultural characteristics.

Many plant pathogens, such as fungi, bacteria, phytoplasmas, viruses, and nematodes can infect grapes, and the resultant diseases can cause substantial losses in production (Pearson et al., Compendium of Grape Diseases, American Phytopathological Society Press (1988)). Among these, viral diseases constitute a major hindrance to profitable growing of grapevines. About 34 viruses have been isolated and characterized from grapevines. The major virus diseases are grouped into: (1) the grapevine degeneration caused by the fanleaf nepovirus, other European nepoviruses, and American nepoviruses, (2) the leafroll complex, and (3) the rugose wood complex (Martelli, ed., Graft Transmissible Diseases of Grapevines, Handbook for Detection and Diagnosis, FAO, UN, Rome, Italy (1993)).

Rugose wood (RW) complex is a term to describe a group of grafttransmissible diseases which are important and widespread on grapevines grown world-wide. Symptoms of RW are characterized by pitting, grooving, or distortion to the woody cylinder of the grapevine scion, rootstock, or both. Based on symptoms developed on different indicator plants after graft inoculation, RW complex can be divided into four components: Kober 5BB stem grooving (KSG), LN 33 stem grooving (LNSG), grapevine corky bark (GCB), and Rupestris stem pitting (RSP) (Martelli, "Rugose Wood Complex," in Graft-Transmissible Diseases of Grapevines, Handbook for Detection and Diagnosis, pp. 45-54, Martelli, ed., Food and Agriculture Organization of the United Nations, Rome, Italy (1993)). Because RW can cause severe decline and death to grapevines (Savino et al., "Rugose Wood Complex of Grapevine: Can Grafting to Vitis Indicators Discriminate Between Diseases?", in Proceedings of the 9th Meetings of the International Council for the Study of Viruses and Virus Diseases of the Grapevine, Anavim, Israel (1989); Credi and Babini, "Effect of Virus and Virus-like Infections on the Growth of Grapevine Rootstocks," Adv. Hort. Sci., 10:95-98 (1996)), it has been included in healthy grapevine detection schemes used in major grapevine growing countries including Italy, France, and the United States.

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RSP was discovered in California in the late 1970s (Prudencio, "M. Sc. Thesis: Comparative Effects of Corky Bark and Rupestris Stem Pitting Diseases on Selected Germplasm Lines of Grapes," University of California, Davis, California, 36 pages (1985); Goheen, "Rupestris Stem Pitting," in Compendium of Grape Diseases, p. 53, Pearson and Goheen, eds., American Phytopathological Society Press, St. Paul, Minnesota, USA (1988) ("Goheen")). The disease was defined by Goheen as follows: after graft inoculation with a chip bud from an infected grapevine, the woody cylinder of the indicator plant Vitis rupestris Scheele St. George ("St. George") develops a narrow strip of small pits extending from the inoculum bud to the root zone. Grafted St. George plants were checked for wood symptoms 2 to 3 years after inoculation. In contrast to GCB, which elicits pitting and grooving on St. George and LN 33, RSP does not produce symptoms on the latter (Goheen, "Rupestris Stem Pitting," in Compendium of Grape Diseases, p. 53, Pearson and Goheen, eds., American Phytopathological Society Press, St. Paul, Minnesota, USA (1988)).

RSP is probably the most common component of the RW complex on grapevines. Surveys in California revealed a high disease incidence in many grapevine cultivars imported from Western Europe and Australia (Goheen, "Rupestris Stem Pitting," in Compendium of Grape Diseases, p. 53, Pearson and Goheen, eds., American Phytopathological Society Press, St. Paul, Minnesota, USA (1988)). An examination of indexing records in California compiled over 23 years revealed RSP infection in 30.5% of 6482 grapevine selections introduced from around the world (Golino and Butler, "A Preliminary Analysis of Grapevine Indexing Records at Davis, California," in Proceedings of the 10th Meeting of the ICVG, pp. 369-72, Rumbos et al., eds., Volos, Greece (1990)). Indexing in New York State showed that 66% of 257 grapevines tested on St. George developed typical small pits below the inoculum bud or around the woody cylinder (Azzam and Gonsalves, Abstract: "Survey of Grapevine Stem-Pitting in New York and Isolation of dsRNA from a Grapevine Selection Infected with Stem Pitting," Phytopathology 78:1568 (1988)). Furthermore, several reports have indicated that RSP is the most frequently detected component of the RW complex in Italy (Borgo and Bonotto, "Rugose Wood Complex of Grapevine in Northeastern Italy: Occurrence of Rupestris Stem Pitting and Kober Stem Grooving," in Extended Abstracts of the 11th Meeting of the International Council for the Study of Viruses and Virus Diseases of the Grapevine (ICVG), pp. 61-62, Gugerli, ed.,

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Montreux, Switzerland (1993); Credi, "Differential Indexing Trials on Grapevine Rugose Wood Syndrome," Extended Abstracts of the 11th Meeting of the International Council for the Study of Viruses and Virus Diseases of the Grapevine (ICVG), p. 63, Gugerh, P., ed., Montreux, Switzerland (1993)).

The effect of RSP on growth, yield, and grapevine quality is not well understood and, thus, subject to debate. The reason for this ambiguity is the absence of a rapid and sensitive diagnostic tool. RSP is the most difficult grapevine disease to diagnose. Serological or molecular methods are not available for diagnosing RSP. Biological indexing on St. George, as described above, has remained the only approach to diagnose RSP. Biological indexing is labor intensive, time consuming (i.e., often requiring up to about three years to obtain results), and, by its very nature, subjective. Moreover, symptoms on St. George can be variable and not exactly as those defined by Goheen. In particular, Credi, "Characterization of Grapevine Rugose Wood Sources from Italy," Plant Disease, 82:1288-92 (1997), recently showed that some RSP infected grapevines induced pitting that is restricted to below the inoculum bud, while others induced pitting around the woody cylinder of inoculated St. George. Thus, the present method of identifying the presence of RSP is not entirely adequate.

The etiology of RSP is unknown. Efforts to isolate virus particles from RSP-infected grapevines and to mechanically transfer the causal virus(es) to 20 herbaceous host plants failed (Azzam and Gonsalves, "Detection of in Grapevines Showing Symptoms of Rupestris Stem Pitting Disease and the Variabilities Encountered," Plant Disease, 75:96-964 (1991)). However, a major dsRNA species of ca. 8.3 kb, accompanied by a smaller dsRNA of ca. 7.6 kb, was consistently isolated from one Pinot Gris and four Pinot Noir clones that had been indexed positive 25 for RSP (Walter and Cameron, "Double-Stranded RNA Isolated from Grapevines Affected by Rupestris Stem Pitting Disease," Am. J. of Enology and Viticulture, 42:175-79 (1991)). In addition, a third dsRNA of ca. 5.5 kb was observed in three clones. Likewise, an apparently similar dsRNA species of ca. 8.0 and 6.7 kbp was 30 isolated from dormant canes of RSP-infected grapevines collected from California, Canada, and New York (Azzam and Gonsalves, "Detection of dsRNA in Grapevines Showing Symptoms of Rupestris Stem Pitting Disease and the Variabilities Encountered," Plant Disease, 75:960-64 (1991)). Six of eight Californian and three of

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five Canadian samples contained these two dsRNA species. However, results of New York samples were not consistent. Among eight RSP infected grapevine selections tested, only one showed these two dsRNAs. Using explants growing in tissue culture as source materials, dsRNA of ca. 359 bp was isolated from 21 of 31 grapevine cultivars, all of which were previously indexed on St. George and considered to be infected with RSP (Monette et al., "Double-Stranded RNA from *Rupestris* Stem Pitting-Affected Grapevines," Vitis, 28:137-44 (1989)).

In view of the serious risk RSP poses to vineyards and the absence of an effective treatment of it, the need to prevent this affliction continues to exist.

Moreover, the absence of a rapid and accurate diagnostic assay prevents proper identification of RSP. The present invention is directed to overcoming these deficiencies in the art.

SUMMARY OF THE INVENTION

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The present invention relates to an isolated protein or polypeptide corresponding to a protein or polypeptide of a RSP virus. The encoding RNA molecule or DNA molecule, in either isolated form or incorporated in an expression system, a host cell, or a transgenic *Vitis* scion or rootstock cultivar, are also disclosed.

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Another aspect of the present invention relates to a method of imparting RSP virus resistance to *Vitis* scion or rootstock cultivars by transforming them with a DNA molecule encoding the protein or polypeptide corresponding to a protein or polypeptide of a RSP virus.

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The present invention also relates to an antibody or binding portion thereof or probe which recognizes proteins or polypeptides of the present invention.

Still another aspect of the present invention relates to diagnostic tests which involve methods for detecting the presence of a RSP virus in a sample. The methods include the use of an antibody or binding portion of the present invention (i.e., in an immunoassay), or a nucleic acid probe obtained from a DNA molecule of the present invention (i.e., in a nucleic acid hybridization assay or gene amplification detection procedure). The antibody or binding portion thereof, or nucleic acid probe, is introduced into contact with the sample, whereby the presence of *Rupestris* stem pitting virus in the sample is detected using an assay system.

The characterization of an RSP virus is particularly desirable because it will allow for the determination of whether the virus is associated to the specific (restricted) or nonspecific (nonrestricted) pitting symptoms of RSP, or to both. Also, RSP virus resistant transgenic variants of the current commercial grape cultivars and rootstocks allows for more complete control of the virus while retaining the varietal characteristics of specifics cultivars. Furthermore, these variants permit control over RSP virus transmitted by infected scions or rootstocks. Moreover, the diagnostic tests offer significant improvement over conventional diagnostic means currently employed, namely, rapid results and greater accuracy.

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BRIEF DESCRIPTION OF THE DRAWINGS

Figure 1 is a photograph of St. George indicators which comparatively display the symptoms of RSP. The St. George indicator (a) has been graft-inoculated with infected bud wood from a grapevine accession, resulting in the indicator displaying pitting below the inoculum bud, as indicated by an arrow. This RSP symptom was defined by Goheen, "Rupestris Stem Pitting," in Compendium of Grape Diseases, p. 53, Pearson and Goheen, eds., American Phytopathological Society Press, St. Paul, Minnesota, USA (1988), which is hereby incorporated by reference. The St. George indicator (b) was not graft-inoculated and represents a normal appearance.

Figures 2A and 2B are photographs which respectively display the results of dsRNA analysis and Northern hybridization for dsRNA. Together the photographs may be used to correlate the dsRNA analysis of Figure 2A with the Northern hybridization (for dsRNA isolated from grapevines indexed positive for *Rupestris* stem pitting (RSP)) of Figure 2B. M. *Hind* III digested lambda DNA maker: lane 1, Aminia; lane 2, Bertille Seyve 5563; lane 3, Canandaigua; lane 4, Colobel 257; lane 5, Couderc 28-112; lane 6, Freedom; lane 7, Grande Glabre; lane 8, M 344-1; lane 9, Joffre; lane 10, Ravat 34; lane 11, Seyval; lane 12, Seyve Vinard 14-287; lane 13, Verdelet; lane 14, Pinot Noir (positive control); lane 15, Verduzzo 233A (negative control for RSP as judged by indexing on St. George); lane 16, insert of clone RSP149. Arrows indicate the position of the 8.7 kb dsRNA. With respect to lane 15 of Figure 2A, the two dsRNA bands are larger or smaller than the 8.7 kb

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dsRNA associated with RSP and they did not hybridize with the RSP specific probe in Northern analysis. Thus, they are not specific to RSP.

Figure 3A is an illustration which depicts the strategy for obtaining the complete nucleotide sequence of RSPaV-1. The overlapping regions of the nucleotide sequences of the sequenced clones and RT-PCR-amplified cDNA fragments are as follows: 52-375 for RSPA/RSP28; 677-1474 for RSP28/RSP3; 3673-3766 for RSP3/RSPB; 4009-4320 for RSPB/RSP94; 5377-5750 for RSP94/RSPC; 5794-6537 for RSPC/RSP95; 6579-6771 for RSPC/RSP140; and 8193-8632 for RSP140/TA5. Figure 3B is an illustration which comparatively depicts the genome structures of RSPaV-1, ASPV, PVM, and PVX. Boxes with the same patterns represent the comparable ORFS.

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Figure 4A is a comparative sequence listing of amino acid sequences of region I (aa 1-372) of RSPaV-1 ORF1 with the corresponding sequences of carlavirus PVM and ASPV. The methyltransferase motif is underlined. Capital letters indicate consensus residues. Figure 4B is a comparative sequence listing of amino acid sequences of region II (aa 1354 to end) of RSPaV-1 ORF1 with the corresponding regions of ASPV and PVM carlavirus. In Figure 4B, the NTP binding motif is underlined at (A) and the GDD containing sequence is underlined at (B). In Figures 4A and 4B, capital letters indicate consensus residues, the symbol * indicates identical amino acid residues between RSPaV-1 and ASPV, and the symbol # indicates identical amino acid residues between RSPaV-1 and PMV.

Figures 5A-D are comparative sequence listings of amino acid sequences for ORF2, ORF3, ORF4, and a C-terminal part of ORF5 (CP) of RSPaV-1, respectively, with ASPV and PVM carlavirus. In Figure 5A, the NTP binding motif, located near the C terminus of ORF2, is underlined. In Figure 5D, the conserved motif (RR/QX--XFDF), located in the central region of the coat proteins and proposed to be involved in the formation of a salt bridge structure, is underlined. In each of the figures, capital letters indicate consensus residues. The symbol * indicates identical amino acid residues between RSPaV-1 and ASPV, and the symbol # indicates identical amino acid residues between RSPaV-1 and PMV. In Figure 5D, numbers which appear in parentheses and precede the sequences indicate the start points of the C-terminal portions of CPs being compared.

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Figure 6A is a comparative sequence listing of DNA nucleotide sequences for the 3' untranslated region (UTR) of RSPaV-1 and ASPV. Figure 6B is a comparative sequence listing of DNA nucleotide sequences for the 3' untranslated region (UTR) of RSPaV-1 and PVM. Clustal method of MegAlign (DNASTAR) was used to generate sequence alignments. The 21 identical consecutive nucleotides between RSPaV-1 and PVM are indicated as shadowed letters.

Figures 7A-B are photographs comparing the results of RT-PCR of grapevines using RSP149 primers (Figure 7A) and Southern blot hybridization of RT-PCR amplified cDNA fragments to RSPaV-1 specific probe (Figure 7B). MMLV-RT (Promega) was used in reverse transcription. *Taq* DNA polymerase (Promega) was used in PCR. For the RT-PCR and Southern blot hybridization: lane 1, Ehrenfelser PM1 (1169-1A1); lane 2, Cabernet franc 147A; lane 3, Chardonnay 80A; lane 4, Refosco 181A; lane 5, Touriga francesa 313; lane 6, 3309C (330-4A1); lane 7, 420A (1483-4A1); lane 8, Chardonnay 83A; lane 9, Malsavia 153A; lane 10, Aragnonex 350; lane 11, Aminia; lane 12, Chardonnay 127; lane 13, Kober 5BB 100; lane 14, Verduzzo 233A; lane 15, *V. riparia*; lane 16, *V. monticola*; lane 17, H₂O.

Figure 8 is a schematic representation of the identical genome organization among RSPaV-1 (the type strain), RSP47-4, and RSP158. The number of amino acid residues of the comparable ORFs (boxes shaded with the same pattern) among these three strains are the same (note: ORF1 and ORF5 of RSP47-4 and RSP158 are incomplete). The comparable ORFs also have high nucleotide and amino acid sequence identities, which are indicated on the bottom. Only the C-terminal portion of the ORF1 of RSPaV-1 is shown in this diagram.

Figure 9 is a comparative alignment of nucleotide sequences of seven other clones with the comparable region of RSPaV-1. Shaded areas indicate identical nucleotide sequences, whereas white boxes represent different nucleotide sequences.

Figure 10 is a schematic representation of a plant transformation vector containing the RSPaV-1 coat protein gene. This vector is designated pGA482G/RSPaV-1CP, which has the double CaMV 35S enhancers, the 35S promoter, the leader sequence of AlMV, and the 35S terminator sequence. RB, right border; LB, left border; Tet, tetracycline resistance gene; and Gent, gentamycin resistance gene.

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DETAILED DESCRIPTION OF THE INVENTION

The present invention relates to isolated DNA molecules encoding for the proteins or polypeptides of a *Rupestris* stem pitting associated virus. Since the nucleotide sequence was derived from cDNA clones of the dsRNA that was associated with RSP, the viral agent has been designated as *Rupestris* stem pitting associated virus ("RSPaV"). RSP is likely caused by one or a number of viral strains. The genome of each RSPaV has a plurality of open reading frames, each containing DNA molecules in accordance with the present invention. The complete genome of one strain has been sequenced and the strain is designated RSPaV-1. Substantial portions of the genomes of two other RSPaV strains have also been sequenced. These strains are designated by their clone names, RSP47-4 and RSP158.

The DNA molecule which constitutes the complete RSPaV-1 genome comprises the nucleotide sequence corresponding to SEQ. ID. No. 1 as follows:

,	CGATAAACAT	AACAACAGAA	TCTGCATTGC	AGTAATATTC	CTTGAATATA	ATTGCAACGC	60
	AATGGCCCTC	TCTTATAGGC	CTGCTGTTGA	AGAGGTGCTC	GCAAAATTCA	CCTCTGATGA	120
	ACAATCCAGG	GTTTCTGCTA	CAGCTCTCAA	GGCATTAGTA	GACTTAGAGG	AAAGTCAGCA	180
	CAATTTGTTC	TCTTTCGCAT	TGCCTGATAG	AAGCAAAGAA	AGGCTGATAT	CTTCTGGCAT	240
	TTACTTAAGT	CCTTACAGTT	TCAGACCCCA	CTCACATCCA	GTTTGTAAAA	CTTTAGAAAA	300
	TCACATTTTG	TACAATGTTT	TACCTAGTTA	TGTTAATAAT	TCATTTTACT	TTGTAGGAAT	360
	CAAGGATTTT	AAGCTGCAGT	TCTTGAAAAG	GAGGAATAAG	GATCTCAGCT	TGGTAGCACT	420
	CATAAATAGG	TTTGTGACAA	GTCGTGATGT	TAGTAGGTAT	GGGTCTGAGT	TCGTTATAAG	480
	TTCTAGTGAC	AAATCAAGTC	AGGTTGTCAG	TAGAAAGGGC	ATTGGTGATT	CTAACACACT	540
	CCGGAGATTG	GTCCCACGTG	TAATTTCCAC	AGGTGCCAGG	AATCTTTTTC	TGCATGATGA	600
	GATTCACTAC	TGGTCAATTA	GTGATCTGAT	CAATTTTTTG	GACGTTGCCA	AGCCAAGCAT	660
	GCTCTTGGCA	ACTGCAGTAA	TCCCTCCAGA	AGTGCTGGTT	GGCTCTCCAG	AGAGTCTTAA	720
	CCCTTGGGCC	TACCAGTATA	AAATCAATGG	CAACCAACTG	CTCTTCGCAC	CAGATGGCAA	780
	CTGGAATGAG	ATGTACTCAC	AACCTTTGTC	: ATGCAGATAC	CTGCTCAAGG	CCAGATCTGT	840
	AGTTCTGCCC	GATGGCTCAC	GCTACTCGGT	TGACATCATT	CACTCAAAAT	TTAGTCACCA	900
	CTTGCTTAGT	TTCACCCCTA	TGGGTAATC	TTTGACTTCA	AACATGCGAT	GTTTTTCTGG	960
	CTTCGATGCA	A ATAGGCATA	AAGATCTTG	A ACCTCTAAGO	CGCGGCATGC	ACAGTTGCTT	1020

CCCAGTACAT	CATGATGTTG	TAACTAAGAT	ATATCTTTAT	TTGAGAACTC	TCAAGAAGCC	1080
AGATAAGGAG	TCTGCCGAGG	CAAAGCTTCG	ACAACTCATA	GAAAAACCCA	CAGGGAGGGA	1140
GATAAAGTTT	ATCGAGGATT	TTTCCTCACT	AGTAATAAAT	TGTGGGAGGA	GTGGCTCTTT	1200
GCTTATGCCC	AACATTTCTA	AGTTGGTCAT	ATCATTCTTT	TGCCGGATGA	TGCCAAATGC	1260
ACTCGCCAGG	CTCTCTTCTA	GCTTTCGAGA	GTGTTCGCTA	GATTCATTTG	TGTACTCACT	1320
TGAGCCCTTT	AATTTTTCCG	TTAATTTAGT	GGATATAACT	CCTGATTTCT	TTGAGCATTT	1380
ATTTCTCTTC	TCCTGCCTAA	ATGAGTTGAT	CGAGGAGGAC	GTTGAAGAGG	TCATGGACAA	1440
TTCTTGGTTT	GGACTTGGGG	ACTTACAATT	CAATCGCCAG	AGGGCCCCGT	TCTTTCTTGG	1500
GTCTTCATAT	TGGCTCAACT	CCAAATTTTC	AGTTGAGCAC	AAGTTTTCAG	GCACCATCAA	1560
TTCTCAAATC	ATGCAAGTTA	TTTTATCTTT	GATCCCATTT	TCTGATGATC	CCACTTTTAG	1620
GCCATCTTCT	ACAGAGGTTA	ACCTTGCACT	ATCAGAGGTT	AAGGCTGCGC	TAGAAGCTAC	1680
TGGGCAGTCA	AAATTGTTCA	GGTTTTTGGT	GGACGACTGT	GCTATGCGTG	AGGTTAGAAG	1740
TTCCTATAAG	GTGGGCCTTT	TTAAGCACAT	AAAAGCCCTC	ACTCATTGCT	TTAATTCTTG	1800
TGGCCTCCAA	TGGTTCCTCC	TTAGGCAAAG	GTCCAACCTC	AAATTTCTGA	AGGACAGGGC	1860 ⁻
ATCGTCCTTT	GCTGATCTTG	ATTGTGAGGT	TATCAAAGTT	TATCAGCTTG	TAACATCACA	1920
GGCAATACTT	CCTGAGGCTC	TGCTTAGCTT	GACCAAAGTC	TTTGTCAGGG	ATTCTGACTC	1980
AAAGGGTGTT	TCCATTCCCA	GATTGGTCTC	GAGAAATGAG	CTAGAGGAAC	TAGCTCACCC	2040
AGCTAATTCA	GCCCTTGAGG	AGCCTCAATC	AGTTGATTGT	AATGCAGGCA	GGGTTCAAGC	2100
AAGCGTTTCA	AGTTCCCAGC	AGCTTGCCGA	CACCCACTCT	CTTGGTAGCG	TTAAGTCATC	2160
AATTGAGACA	GCTAACAAGG	CTTTTAACTT	GGAGGAGCTA	AGGATCATGA	A TTAGAGTCTT	2220
GCCGGAGGAT	TTTAACTGGG	TGGCGAAGAA	CATTGGTTTT	AAAGACAGGG	TGAGAGGCAG	2280
GGGTGCATCA	TTCTTCTCAA	AACCAGGAA1	TTCATGTCAT	AGTTACAATO	GTGGGAGCCA	2340
CACAAGCTTA	GGGTGGCCAA	AGTTCATGGA	TCAGATTCTA	AGCTCCACTO	G GTGGACGTAA	2400
TTACTACAAT	TCATGCCTGG	CTCAGATCT/	TGAGGAAAAI	TCAAAATTG	CTCTTCATAA	2460
GGATGATGAG	AGTTGCTATG	AAATTGGGC	CAAAGTTTT	ACTGTTAAT	TAATCGGCTC	2520
AGCAACTTTC	CACTATTAGTA	AGTCGCGAA	A TTTGGTTGG	GGTAATCAT	T GCAGCCTGAC	2580
AATTGGGCCA	AATGAGTTTT	TCGAAATGC	C TAGGGGCATO	CAATGCAAT	T ACTTCCATGG	2640
GGTTTCCAAT	TGTACGCCAG	GGCGGGTAT	C GCTGACCTT	r aggcgccaa	A AGTTGGAAGA	2700
TGATGATTT	ATCTTCATA	ATCCACAGG	r GCCCATTGAC	G CTCAATCAT	G AAAAGCTTGA	2760
CCGAAGTATO	TGGCAGATGG	GCCTTCATG	G AATTAAGAA	A TCTATTTCT	A TGAATGGCAC	2820

GAGTTTTACC TCAGACCTAT GCTCTTGTTT CTCTTGCCAC AACTTTCATA AATTCAAGGA	2880
TCTCATCAAT AACTTGAGAT TGGCCCTAGG AGCACAAGGG CTAGGTCAGT GTGACAGGGT	2940
TGTGTTTGCA ACAACAGGTC CTGGTCTATC TAAGGTTTTA GAAATGCCTC GGAGCAAAAA	3000
GCAATCAATT TTGGTTCTTG AAGGTGCCCT ATCCATAGAA ACAGATTATG GTCCAAAAGT	3060
CCTGGGGTCT TTTGAAGTTT TCAAAGGGGA CTTTCACATT AAGAAGATGG AGGAAGGTTC	3120
AATTTTTGTA ATAACGTACA AGGCCCCAAT TAGATCCACT GGCAGGTTGA GGGTTCACAG	3180
TTCAGAATGC TCATTTTCCG GATCCAAAGA GGTATTGCTA GGCTGCCAGA TTGAGGCATG	3240
TGCTGATTAT GATATTGATG ATTTTAACAC TTTCTCTGTG CCTGGTGATG GCAATTGCTT	3300
TTGGCATTCT GTTGGTTTTT TACTTAGCAC TGATGGACTT GCCCTAAAGG CCGGTATTCG	3360
ATCTTTCGTG GAGAGTGAGC GCTTGGTAAG TCCAGATCTT TCAGCCCCAG CAATTTCTAA	3420
ACAATTGGAA GAGAATGCTT ATGCCGAGAA TGAGATGATC GCATTATTCT GCATTCGGCA	3480
CCACGTAAGG CCTATAGTGA TCACACCAGA ATATGAAGTT AGTTGGAAAT TCGGGGAAGG	3540
TGAGTGGCCC CTATGTGGAA TTCTTTGCCT TAAATCAAAT CACTTCCAAC CATGCGCCCC	3600
ACTGAATGGT TGCATGATCA CAGCCATTGC TTCAGCACTT GGAAGGCGTG AAGTTGATGT	3660
GTTAAATTAT CTGTGTAGAC CCAGCACTAA TCATATTTTT GAGGAGCTTT GTCAGGGAGC	3720
GGGCCTTAAC ATGATGTATT TAGCTGAAGC TTTTGAGGCC TTTGACATTT GCGCTAAAT	3780
TGATATAAAT GGAGAGATTG AAGTGATTAA TCCGTGTGGT AAAATTTCTG CATTGTTTG	A ~ 3840
CATAACTAAT GAGCACATAA GGCATGTTGA GAAAATAGGT AATGGCCCTC AGAGCATAA	A 3900
AGTGGATGAA TTGCGGAAGG TCAAGCGATC CGCCCTCGAT TTCCTTTCAA TGAATGGGT	C 3960
TAAAATAACC TACTTCCCAA GCTTTGAGCG GGCTGAAAAG TTGCAAGGAT GTTTGCTAG	G 4020
GGGCCTAACT GGCGTTATAA GTGATGAGAA GTTCAGTGAT GCAAAACCTT GGCTTTCTG	G 4080
TATATCTACT ACTGATATTA AGCCAAGGGA ATTGACTGTC GTGCTTGGTA CATTTGGGG	C 4140
TGGGAAGAGT TTCTTGTACA AGAGTTTCAT GAAAAGGTCT GAGGGTAAAT TCGTAACCT	T 4200
TGTTTCTCCC AGACGTGCTT TAGCAAATTC AATCAAAAAT GATCTTGAAA TGGATGATA	AG 4260
CTGCAAAGTT GCTAAAGCAG GTAGGTCAAA GAAGGAAGGG TGGGATGTAG TAACTTTTC	SA 4320
GGTTTTCCTT AGAAAAGTTG CAGGATTGAA GGCTGGCCAC TGTGTGATTT TTGATGAG	GT 4380
CCAGTTGTTT CCTCCTGGAT ACATCGATCT ATGCTTGCTT ATTATACGTA GTGATGCT	TT 4440
CATTICACTI GCTGGTGATC CATGTCAAAG CACATATGAC TCGCAAAAGG ATCGGGCA	AT 4500
TTTGGGCGCT GAGCAGAGTG ACATACTTAG ACTGCTTGAG GGCAAAACGT ATAGGTAT	AA 4560
CATAGAAAGC AGGAGGTTTG TGAACCCAAT GTTCGAATCA AGACTGCCAT GTCACTTC	AA 4620

AAAGGGCTCG	ATGACTGCCG	CTTTCGCTGA	TTATGCAATC	TTCCATAATA	TGCATGACTT	4680
TCTCCTGGCG	AGGTCAAAAG	GTCCCTTGGA	TGCCGTTTTG	GTTTCCAGTT	TTGAGGAGAA	4740
AAAGATAGTC	CAGTCCTACT	TTGGAATGAA	ACAGCTCACA	CTCACATTTG	GTGAATCAAC	4800
TGGGTTGAAT	TTCAAAAATG	GGGGAATTCT	CATATCACAT	GATTCCTTTC	ACACAGATGA	4860
TCGGCGGTGG	CTTACTGCTT	TATCTCGCTT	CAGCCACAAT	TTGGATTTGG	TGAACATCAC	4920
AGGTCTGAGG	GTGGAAAGTT	TTCTCTCGCA	CTTTGCTGGC	AAACCCCTCT	ACCATTTTTT	4980
AACAGCCAAA	AGTGGGGAGA	ATGTCATACG	AGATTTGCTC	CCAGGTGAGC	CTAACTTCTT	5040
CAGTGGCTTT	AACGTTAGCA	TTGGAAAGAA	TGAAGGTGTT	AGGGAGGAGA	AGTTATGTGG	5100
TGACCCATGG	TTAAAAGTTA	TGCTTTTCCT	GGGTCAAGAT	GAGGATTGTG	AAGTTGAAGA	5160
GATGGAGTCA	GAATGCTCAA	ATGAAGAATG	GTTTAAAACC	CACATCCCCT	TGAGTAATCT	5220
GGAGTCAACC	AGGGCCAGGT	GGGTGGGTAA	AATGGCCTTG	AAAGAGTATC	GGGAGGTGCG	5280
TTGTGGTTAT	GAAATGACTC	AACAATTCTT	TGATGAGCAT	AGGGGTGGAA	CTGGTGAGCA	5340
ACTGAGCAAT	GCATGTGAGA	GGTTTGAAAG	CATTTACCCA	AGGCATAAAG	GAAATGATTC	5400
AATAACCTTC	CTCATGGCTG	TCCGAAAGCG	TCTCAAATTT	TCGAAGCCCC	AGGTTGAAGC	5460
TGCCAAACTG	AGGCGGGCCA	AACGATATGG	GAAATTCTTA	TTAGATTCTT	TCCTATCCAA	5520
AATCCCATTG	AAAGCCAGTC	ATAATTCCAT	CATGTTTCAT	GAAGCGGTAC	AGGAGTTTGA	5580
GGCGAAGAAG	GCTAGTAAGA	GTGCAGCAAC	TATAGAGAAT	CATGCAGGTA	GGTCATGCAG	5640
GGATTGGTTA	TTAGATGTTG	CTCTGATTTT	TATGAAGTCA	CAACACTGTA	CTAAATTTGA	5700
CAACAGGCTT	AGAGTAGCTA	AAGCTGGGCA	AACCCTTGCT	TGCTTCCAAC	ATGCTGTTCT	5760
GGTTCGCTTT	GCACCCTATA	TGAGATACAT	TGAGAAAAAG	CTAATGCAAC	CTCTGAAGCC	5820
TAACTTCTAC	ATCCATTCAG	GGAAAGGTCI	GACGAGCTGA	ACGAGTGGG	r cagaactaga	5880
GGATTCACTG	GAATTTGCAC	AGAATCAGAG	TACGAAGCCI	TTGATGCTT	CCAAGACCAC	5940
TTCATCCTAG	CATTCGAATT	GCAGATAATO	AAATTTTTGG	GGTTACCTG	A AGATTTAATT	6000
TTGGACTATO	AATTCATAAA	AATTCATTT	G GGATCAAAG	TCGGATCAT	r CTCTATAATG	6060
AGGTTTACT	GGGAGGCCAG	CACATTTCT	TTTAACACT	A TGGCTAACA	T GTTGTTCACC	6120
TTTCTGAGGT	CACGAACTAAC	AGGCTCTGAG	G TCAATAGCA	TTGCAGGTG	A TGACATGTGT	6180
GCTAATCGA	A GGTTGCGGCT	TAAAACAGA	G CATGAGGGT	r ttctgaaca	T GATTTGCCTT	6240
AAGGCCAAGG	TTCAGTTTG	TTCCAATCC	C ACATTCTGC	G GATGGTGTT	T ATTTAAGGAA	6300
GGGATCTTC	A AGAAGCCTC	ATTAATCTG	G GAGCGGATA	T GCATTGCTA	G GGAGATGGGC	6360
AACCTGGAGA	A ATTGTATTG	A CAATTATGO	G ATAGAGGTC	T CCTATGCAT	A CCGACTGGGA	6420

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GAGCTAGCCA	TTGAAATGAT	GACCGAGGAA	GAAGTGGAGG	CCCATTATAA	TTGTGTTAGA	6480
TTCTTGGTCA	GGAACAAGCA	TAAGATGAGA	TGCTCAATTT	CAGGCCTATT	TGAAGCTATT	6540
GATTAGGCCT	TAAGTATTTG	GCATTATTTG	AGTATTATGA	ATAATTTAGT	TAAAGCATTG	6600
TCAGCATTTG	AGTTTGTAGG	TGTTTTCAGT	GTGCTTAAAT	TTCCAGTAGT	CATTCATAGT	6660
GTGCCTGGTA	GTGGTAAAAG	TAGTTTAATA	AGGGAGCTAA	TTTCCGAGGA	TGAGAATTTC	6720
ATAGCTTTCA	CAGCAGGTGT	TCCAGACAGC	CCTAATCTCA	CAGGAAGGTA	CATTAAGCCT	6780
TATTCTCCAG	GGTGTGCAGT	GCCAGGGAAA	GTTAATATAC	TTGATGAGTA	CTTGTCCGTC	6840
CAAGATTTTT	CAGGTTTTGA	TGTGCTGTTC	TCGGACCCAT	ACCAAAACAT	CAGCATTCCT	6900
AAAGAGGCAC	ATTTCATCAA	GTCAAAAACT	TGTAGGTTTG	GCGTGAATAC	TTGCAAATAT	6960
CTTTCCTCCT	TCGGTTTTAA	GGTTAGCAGT	GACGGTTTGG	ACAAAGTCAT	TGTGGGGTCG	7020
CCTTTTACAC	TAGATGTTGA	AGGGGTGCTA	ATATGCTTTG	GTAAGGAGGC	AGTGGATCTC	7080
GCTGTTGCGC	ACAACTCTGA	ATTCAAATTA	CCTTGTGAAG	TTAGAGGTTC	AACTTTTAAC	7140
GTCGTAACTC	TTTTGAAATC	AAGAGATCCA	ACCCCAGAGG	ATAGGCACTG	GTTTTACATT	7200
GCTGCTACAA	GACACAGGGA	GAAATTGATA	ATCATGCAGT	AAGATGCCTT	TTCAGCAGCC	7260
TGCGAATTGG	GCAAAAACCA	TAACTCCATT	GACAGTTGGC	TTGGGCATTG	GGCTTGTGCT	7320
GCATTTTCT	G AGGAAGTCAA	ATCTACCTTA	TTCAGGGGAC	AACATCCATC	AATTCCCTCA	7380
CGGTGGGCGT	TACAGGGACG	GTACAAAAAG	TATAACTTAG	TGTGGTCCAA	AGCAATCCTT	7440
CCCCAGCTC	GGGATATTCC	GCCAATCTG#	A GAATTTTGT	G CCCTTAATGO	TTGTCATAGG	7500
TCTAATCGC	A TTCATACATO	TATTGTCTGT	TTGGAATTC	r ggtcttggt <i>i</i>	GGAATTGTAA	7560
TTGCCATCC	A AATCCTTGC	T CATGTAGACA	A GCAGTAGTG	G CAACCACCA	A GGTTGCTTCA	7620
TTAGGGCCA	C TGGAGAGTC	A ATTTTGATT	AAAACTGCG	G CCCAAGTGA	GCCCTTGCAT	7680
CCACTGTGA	A GGAGGTGCT	G GGAGGTTTG	A AGGCTTTAG	G GGTTAGCCG	r GCTGTTGAAG	7740
AAATTGATT	A TCATTGTTA	A ATTGGCTGA	A TGGCAAGTC	A AATTGGGAA	A CTCCCCGGTG	7800
AATCAAATG.	A GGCTTTTGA	A GCCCGGCTA	A AATCGCTGG	A GTTAGCTAG	A GCTCAAAAGC	7860
AGCCGGAAG	G TTCTAATGC	A CCACCTACT	C TCAGTGGCA	T TCTTGCCAA	A CGCAAGAGGA	7920
TTATAGAGA	A TGCACTTTC	A AAGACGGTG	G ACATGAGGG	A GGTTTTGAA	A CACGAAACGG	7980
TGGTGATTT	C CCCAAATGT	C ATGGATGAA	G GTGCAATAG	A CGAGCTGAT	T CGTGCATTTG	8040
GTGAATCTG	G CATAGCTGA	A AGCGTGCAA	T TTGATGTG	CATAGATAT	A GCACGTCACT	8100
GCTCTGATG	T TGGTAGCTC	C CAGAGGTCA	A CCCTGATTO	GG CAAGAGTCC	A TTTTGTGACC	8160
TAAACAGAT	C AGAAATAGO	T GGGATTATA	A GGGAGGTG	AC CACATTACO	T AGATTTTGCA	8220

TGTACTATGC	AAAAATCGTG	TGGAACATCC	ATCTGGAGAC	GGGGATACCA	CCAGCTAACT	8280
GGGCCAAGAA	AGGATTTAAT	GAGAATGAAA	AGTTTGCAGC	CTTTGATTTT	TTCTTGGGAG	8340
TCACAGATGA	GAGTGCGCTT	GAACCAAAGG	GTGGAATTAA	AAGAGCTCCA	ACGAAAGCTG	8400
AGATGGTTGC	TAATATCGCC	TCTTTTGAGG	TTCAAGTGCT	CAGACAAGCT	ATGGCTGAAG	8460
GCAAGCGGAG	TTCCAACCTT	GGAGAGATTA	GTGGTGGAAC	GGCTGGTGCA	CTCATCAACA	8520
ACCCCTTTTC	AAATGTTACA	CATGAATGAG	GATGACGAAG	TCAGCGACAA	TTCCGCAGTC	8580
CAATAATTCC	CCGATTTCAA	GGCTGGGTTA	AGCCTGTTCG	CTGGAATACC	GTACTAATAG	8640
TATTCCCTTT	CCATGCTAAA	TCCTATTTAA	TATATAAGGT	GTGGAAAGTA	AAAGAAGATT	8700
TGGTGTGTTT	TTATAGTTTT	CATTCAAAAA	АААААААА	AAA		8743

The DNA molecule of SEQ. ID. No. 1 contains at least five open reading frames (e.g., ORF1-ORF5), each of which encodes a particular protein or polypeptide of RSPaV-1, and a 3' untranscribed region downstream of ORF5.

Another DNA molecule of the present invention (RSPaV-1 ORF1) includes nucleotides 62-6547 of SEQ. ID. No. 1. The DNA molecule of RSPaV-1 ORF1 encodes for a RSPaV-1 replicase and comprises a nucleotide sequence corresponding to SEQ. ID. No. 2 as follows:

ATGGCCCTCT CTTATAGGCC TGCTGTTGAA GAGGTGCTCG CAAAATTCAC CTCTGATGAA 60 CAATCCAGGG TTTCTGCTAC AGCTCTCAAG GCATTAGTAG ACTTAGAGGA AAGTCAGCAC 120 AATTTGTTCT CTTTCGCATT GCCTGATAGA AGCAAAGAAA GGCTGATATC TTCTGGCATT 180 TACTTAAGTC CTTACAGTTT CAGACCCCAC TCACATCCAG TTTGTAAAAC TTTAGAAAAT 240 CACATTTTGT ACAATGTTTT ACCTAGTTAT GTTAATAATT CATTTTACTT TGTAGGAATC 300 AAGGATTTTA AGCTGCAGTT CTTGAAAAGG AGGAATAAGG ATCTCAGCTT GGTAGCACTC 360 ATAAATAGGT TTGTGACAAG TCGTGATGTT AGTAGGTATG GGTCTGAGTT CGTTATAAGT 420 TCTAGTGACA AATCAAGTCA GGTTGTCAGT AGAAAGGGCA TTGGTGATTC TAACACACTC 480 CGGAGATTGG TCCCACGTGT AATTTCCACA GGTGCCAGGA ATCTTTTTCT GCATGATGAG 540 ATTCACTACT GGTCAATTAG TGATCTGATC AATTTTTTGG ACGTTGCCAA GCCAAGCATG 600 CTCTTGGCAA CTGCAGTAAT CCCTCCAGAA GTGCTGGTTG GCTCTCCAGA GAGTCTTAAC 660 CCTTGGGCCT ACCAGTATAA AATCAATGGC AACCAACTGC TCTTCGCACC AGATGGCAAC 720 TGGAATGAGA TGTACTCACA ACCTTTGTCA TGCAGATACC TGCTCAAGGC CAGATCTGTA 780

GTTCTGCCCG ATGGCTCACG	CTACTCGGTT	GACATCATTC	ACTCAAAATT	TAGTCACÇAC	840
TTGCTTAGTT TCACCCCTAT	GGGTAATCTT	TTGACTTCAA	ACATGCGATG	TTTTTCTGGC	900
TTCGATGCAA TAGGCATAAA	AGATCTTGAA	CCTCTAAGCC	GCGGCATGCA	CAGTTGCTTC	960
CCAGTACATC ATGATGTTGT	AACTAAGATA	TATCTTTATT	TGAGAACTCT	CAAGAAGCCA	1020
GATAAGGAGT CTGCCGAGGG	CAAAGCTTCGA	CAACTCATAG	AAAAACCCAC	AGGGAGGAG	1080
ATAAAGTTTA TCGAGGATTT	TTCCTCACTA	GTAATAAATT	GTGGGAGGAG	TGGCTCTTTG	1140
CTTATGCCCA ACATTTCTA	A GTTGGTCATA	TCATTCTTTT	GCCGGATGAT	GCCAAATGCA	1200
CTCGCCAGGC TCTCTTCTAG	G CTTTCGAGAG	TGTTCGCTAG	ATTCATTTGT	GTACTCACTT	1260
GAGCCCTTTA ATTTTTCCG	TAATTTAGTG	GATATAACTC	CTGATTTCTT	TGAGCATTTA	1320
TTTCTCTTCT CCTGCCTAA	A TGAGTTGATO	GAGGAGGACG	TTGAAGAGGT	CATGGACAAT	1380
TCTTGGTTTG GACTTGGGG	A CTTACAATTO	AATCGCCAGA	GGGCCCCGTT	CTTTCTTGGG	1440
TCTTCATATT GGCTCAACT	C CAAATTTTCA	GTTGAGCACA	AGTTTTCAGG	CACCATCAAT	1500
TCTCAAATCA TGCAAGTTA	T TTTATCTTTC	ATCCCATTTT	CTGATGATCC	CACTTTTAGG	1560
CCATCTTCTA CAGAGGTTA	A CCTTGCACT	A TCAGAGGTTA	AGGCTGCGCT	AGAAGCTACT	1620
GGGCAGTCAA AATTGTTCA	G GTTTTTGGT	GACGACTGTG	CTATGCGTGA	GGTTAGAAGT	1680
TCCTATAAGG TGGGCCTTT	T TAAGCACATA	A AAAGCCCTCA	CTCATTGCTI	TAATTCTTGT	1740
GGCCTCCAAT GGTTCCTCC	T TAGGCAAAG	G TCCAACCTCA	AATTTCTGAA	GGACAGGGCA	1800
TCGTCCTTTG CTGATCTTG	A TTGTGAGGT	r atcaaagttī	ATCAGCTTG	AACATCACAG	1860
GCAATACTTC CTGAGGCTC	T GCTTAGCTT	G ACCAAAGTCI	TTGTCAGGG	A TTCTGACTCA	1920
AAGGGTGTTT CCATTCCCA	G ATTGGTCTC	G AGAAATGAGO	TAGAGGAAC	T AGCTCACCCA	1980
GCTAATTCAG CCCTTGAGG	A GCCTCAATC	A GTTGATTGT	ATGCAGGCA	G GGTTCAAGCA	2040
AGCGTTTCAA GTTCCCAGC	CA GCTTGCCGA	C ACCCACTCT	TTGGTAGCG	I TAAGTCATCA	2100
ATTGAGACAG CTAACAAG	C TTTTAACTT	G GAGGAGCTA	A GGATCATGA	T TAGAGTCTTG	2160
CCGGAGGATT TTAACTGGG	T GGCGAAGAA	C ATTGGTTTT	A AAGACAGGC	T GAGAGGCAGG	2220
GGTGCATCAT TCTTCTCA	AA ACCAGGAAT	T TCATGTCAT	A GTTACAATG	G TGGGAGCCAC	2280
ACAAGCTTAG GGTGGCCA	AA GTTCATGGA	T CAGATTCTA	A GCTCCACTG	G TGGACGTAAT	2340
TACTACAATT CATGCCTG	GC TCAGATCTA	T GAGGAAAAT	T CAAAATTGG	C TCTTCATAAG	2400
GATGATGAGA GTTGCTAT	GA AATTGGGCA	AC AAAGTTTTG	A CTGTTAATT	T AATCGGCTCA	2460
GCAACTTTCA CTATTAGT	AA GTCGCGAA	AT TTGGTTGGG	G GTAATCATT	G CAGCCTGACA	2520
ATTGGGCCAA ATGAGTTT	TT CGAAATGC	CT AGGGGCATG	C AATGCAATI	TA CTTCCATGGG	2580

GTTTCCAATT	GTACGCCAGG	GCGGGTATCG	CTGACCTTTA	GGCGCCAAAA	GTTGGAAGAT	2640
GATGATTTGA	TCTTCATAAA	TCCACAGGTG	CCCATTGAGC	TCAATCATGA	AAAGCTTGAC	2700
CGAAGTATGT	GGCAGATGGG	CCTTCATGGA	ATTAAGAAAT	CTATTTCTAT	GAATGGCACG	2760
AGTTTTACCT	CAGACCTATG	CTCTTGTTTC	TCTTGCCACA	ACTTTCATAA	ATTCAAGGAT	2820
CTCATCAATA	ACTTGAGATT	GGCCCTAGGA	GCACAAGGGC	TAGGTCAGTG	TGACAGGGTT	2880
GTGTTTGCAA	CAACAGGTCC	TGGTCTATCT	AAGGTTTTAG	AAATGCCTCG	GAGCAAAAAG	2940
CAATCAATTT	TGGTTCTTGA	AGGTGCCCTA	TCCATAGAAA	CAGATTATGG	TCCAAAAGTC	3000
CTGGGGTCTT	TTGAAGTTTT	CAAAGGGGAC	TTTCACATTA	AGAAGATGGA	GGAAGGTTCA	3060
ATTTTTGTAA	TAACGTACAA	GGCCCCAATT	AGATCCACTG	GCAGGTTGAG	GGTTCACAGT	3120
TCAGAATGCT	CATTTTCCGG	ATCCAAAGAG	GTATTGCTAG	GCTGCCAGAT	TGAGGCATGT	3180
GCTGATTATG	ATATTGATGA	TTTTAACACT	TTCTCTGTGC	CTGGTGATGG	CAATTGCTTT	3240
TGGCATTCTG	TTGGTTTTTT	ACTTAGCACT	GATGGACTTG	CCCTAAAGGC	CGGTATTCGA	3300
TCTTTCGTGG	AGAGTGAGCG	CTTGGTAAGT	CCAGATCTTT	CAGCCCCAGC	AATTTCTAAA	3360
CAATTGGAAG	AGAATGCTTA	TGCCGAGAAT	GAGATGATCG	CATTATTCTG	CATTCGGCAC	3420
CACGTAAGGC	CTATAGTGAT	CACACCAGAA	TATGAAGTTA	GTTGGAAATT	CGGGGAAGGT	3480
GAGTGGCCCC	TATGTGGAAT	TCTTTGCCTT	AAATCAAATC	ACTTCCAACC	ATGCGCCCCA	3540
CTGAATGGTT	GCATGATCAC	AGCCATTGCT	TCAGCACTTG	GAAGGCGTGA	AGTTGATGTG	3600
TTAAATTATC	TGTGTAGACC	CAGCACTAAT	CATATTTTG	AGGAGCTTTG	TCAGGGAGGG	3660
GGCCTTAACA	TGATGTATTT	AGCTGAAGCT	TTTGAGGCCT	TTGACATTTG	CGCTAAATGT	3720
GATATAAATG	GAGAGATTGA	AGTGATTAAT	CCGTGTGGTA	AAATTTCTGC	ATTGTTTGAC	3780
ATAACTAATG	AGCACATAAG	GCATGTTGAG	S AAAATAGGTA	ATGGCCCTCA	GAGCATAAAA	3840
GTGGATGAAT	TGCGGAAGGT	CAAGCGATC	GCCCTCGATT	TCCTTTCAAT	GAATGGGTCT	3900
AAAATAACCT	ACTTCCCAAG	CTTTGAGCG	GCTGAAAAGT	TGCAAGGAT	TTTGCTAGGG	3960
GGCCTAACTG	GCGTTATAAG	TGATGAGAA	G TTCAGTGAT	CAAAACCTT	GCTTTCTGGT	4020
ATATCTACTA	CTGATATTA	GCCAAGGGA	A TTGACTGTC	G TGCTTGGTA	CATTTGGGGCT	4080
GGGAAGAGTT	TCTTGTACA	GAGTTTCAT	G AAAAGGTCT	G AGGGTAAAT	r cgtaaccttt	4140
GTTTCTCCCA	A GACGTGCTT	AGCAAATTC	A ATCAAAAAT	G ATCTTGAAA'	r GGATGATAGC	4200
TGCAAAGTTG	CTAAAGCAGG	TAGGTCAAA	G AAGGAAGGG	r gggatgtag	T AACTTTTGAG	4260
GTTTTCCTT	A GAAAAGTTGO	CAGGATTGAA	G GCTGGCCAC	T GTGTGATTT	T TGATGAGGTC	4320
CAGTTGTTTC	CTCCTGGAT	A CATCGATCT	A TGCTTGCTT	A TTATACGTA	G TGATGCTTTC	4380

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ATTTCACTTG	CTGGTGATCC	ATGTCAAAGC	ACATATGACT	CGCAAAAGGA	TCGGGCAATT	4440
TTGGGCGCTG	AGCAGAGTGA	CATACTTAGA	CTGCTTGAGG	GCAAAACGTA	TAGGTATAAC	4500
ATAGAAAGCA	GGAGGTTTGT	GAACCCAATG	TTCGAATCAA	GACTGCCATG	TCACTTCAAA	4560
AAGGGCTCGA	TGACTGCCGC	TTTCGCTGAT	TATGCAATCT	TCCATAATAT	GCATGACTTT	4620
CTCCTGGCGA	GGTCAAAAGG	TCCCTTGGAT	GCCGTTTTGG	TTTCCAGTTT	TGAGGAGAAA	4680
AAGATAGTCC	AGTCCTACTT	TGGAATGAAA	CAGCTCACAC	TCACATTTGG	TGAATCAACT	4740
GGGTTGAATT	TCAAAAATGG	GGGAATTCTC	ATATCACATG	ATTCCTTTCA	CACAGATGAT	4800
CGGCGGTGGC	TTACTGCTTT	ATCTCGCTTC	AGCCACAATT	TGGATTTGGT	GAACATCACA	4860
GGTCTGAGGG	TGGAAAGTTT	TCTCTCGCAC	TTTGCTGGCA	AACCCCTCTA	CCATTTTTTA	4920
ACAGCCAAAA	GTGGGGAGAA	TGTCATACGA	GATTTGCTCC	CAGGTGAGCC	TAACTTCTTC	4980
AGTGGCTTTA	ACGTTAGCAT	TGGAAAGAAT	GAAGGTGTTA	GGGAGGAGAA	GTTATGTGGT	5040
GACCCATGGT	TAAAAGTTAT	GCTTTTCCTG	GGTCAAGATG	AGGATTGTGA	AGTTGAAGAG	5100
ATGGAGTCAG	AATGCTCAAA	TGAAGAATGG	TTTAAAACCC	ACATCCCCTT	GAGTAATCTG	5160
GAGTCAACCA	GGGCCAGGTG	GGTGGGTAAA	ATGGCCTTGA	AAGAGTATCG	GGAGGTGCGT	5220
TGTGGTTATG	AAATGACTCA	ACAATTCTTT	GATGAGCATA	GGGGTGGAAC	TGGTGAGCAA	5280
CTGAGCAATG	CATGTGAGAG	GTTTGAAAGC	C ATTTACCCAA	GGCATAAAGG	AAATGATTCA	5340
ATAACCTTCC	: TCATGGCTGT	CCGAAAGCG	r ctcaaattt	CGAAGCCCCA	GGTTGAAGCT	5400
GCCAAACTGA	GGCGGGCCA	ACCATATGG	G AAATTCTTAI	TAGATTCTTI	CCTATCCAAA	5460
ATCCCATTG	AAGCCAGTC	A TAATTCCATO	C ATGTTTCATO	AAGCGGTACA	A GGAGTTTGAG	5520
GCGAAGAAG	CTAGTAAGA	G TGCAGCAAC	T ATAGAGAATO	ATGCAGGTAG	GTCATGCAGG	5580
GATTGGTTA:	TAGATGTTG	C TCTGATTTT	T ATGAAGTCA	AACACTGTAG	TAAATTTGAC	5640
AACAGGCTT	A GAGTAGCTA	A AGCTGGGCA	A ACCCTTGCT	GCTTCCAAC	A TGCTGTTCTG	5700
GTTCGCTTT	G CACCCTATA	r gagatacat	T GAGAAAAAG(C TAATGCAAG	C TCTGAAGCCT	5760
AACTTCTAC	A TCCATTCAG	g gaaaggtct	G ACGAGCTGA	A CGAGTGGGT	C AGAACTAGAG	5820
GATTCACTG	G AATTTGCAC	A GAATCAGAC	T ACGAAGCCT	TGATGCTTC	C CAAGACCACT	5880
TCATCCTAG	C ATTCGAATT	G CAGATAATG	A AATTTTTGG	G GTTACCTGA	A GATTTAATTT	5940
TGGACTATG	A ATTCATAAA	A ATTCATTTG	G GATCAAAGC	T CGGATCATT	C TCTATAATGA	6000
GGTTTACTG	G GGAGGCCAG	C ACATTTCTG	ст ттаасаста	T GGCTAACAT	G TTGTTCACCT	6060
TTCTGAGGT	A CGAACTAAC	A GGCTCTGAG	ST CAATAGCAT	T TGCAGGTGA	T GACATGTGTG	6120
CTAATCGAA	G GTTGCGGCT	T AAAACAGAG	C ATGAGGGTT	T TCTGAACAT	G ATTTGCCTTA	6180

ATTAG					6485
TCTTGGTCAG GAACAAGO	CAT AAGATGAGAT	GCTCAATTTC	AGGCCTATTT	GAAGCTATTG	6480
AGCTAGCCAT TGAAATGA	TG ACCGAGGAAG	AAGTGGAGGC	CCATTATAAT	TGTGTTAGAT	6420
ACCTGGAGAA TTGTATTG	SAC AATTATGCGA	TAGAGGTCTC	CTATGCATAC	CGACTGGGAG	6360
GGATCTTCAA GAAGCCTC	CAA TTAATCTGGG	AGCGGATATG	CATTGCTAGG	GAGATGGGCA	6300
AGGCCAAGGT TCAGTTTG	TT TCCAATCCCA	CATTCTGCGG	ATGGTGTTTA	TTTAAGGAAG	6240

The RSPaV-1 replicase has a deduced amino acid sequence corresponding to SEQ. ID. No. 3 as follows:

Met 1	Ala	Leu	Ser	Tyr 5	Arg	Pro	Ala	Val	Glu 10	Glu	Val	Leu	Ala	Lys 15	Phe
Thr	Ser	Asp	Glu 20	Gln	Ser	Arg	Val	Ser 25	Ala	Thr	Ala	Leu	Lys 30	Ala	Leu
Val	Asp	Leu 35	Glu	Glu	Ser	Gln	His 40	Asn	Leu	Phe	Ser	Phe 45	Ala	Leu	Pro
Asp	Arg 50	Ser	Lys	Glu	Arg	Leu 55	Ile	Ser	Ser	Gly	Ile 60	Tyr	Leu	Ser	Pro
Tyr 65	Ser	Phe	Arg	Pro	His 70	Ser	His	Pro	Val	Cys 75	Lys	Thr	Leu	Glu	Asn 80
His	Ile	Leu	Tyr	Asn 85	Val	Leu	Pro	Ser	Tyr 90	Val	Asn	Asn	Ser	Phe 95	Tyr
Phe	Val	Gly	Ile 100	Lys	Asp	Phe	Lys	Leu 105	Gln	Phe	Leu	Lys	Arg 110	Arg	Asn
Lys	Asp	Leu 115	Ser	Leu	Val	Ala	Leu 120	Ile	Asn	Arg	Phe	Val 125	Thr	Ser	Arg
Asp	Val 130	Ser	Arg	Tyr	Gly	Ser 135	Glu	Phe	Val	Ile	Ser 140	Ser	Ser	Asp	Lys
Ser 145	Ser	Gln	Val	Val	Ser 150	Arg	Lys	Gly	Ile	Gly 155		Ser	Asn	Thr	Leu 160
Arg	Arg	Leu	Val	Pro 165	_	Val	Ile	Ser	Thr 170	_	Ala	Arg	Asn	Leu 175	Phe
Leu	His	Asp	Glu 180		His	Tyr	Trp	Ser 185		Ser	Asp	Leu	11e 190		Phe
Leu	Asp	Val 195		Lys	Pro	Ser	Met 200		Lev	a Ala	Thr	Ala 205		Ile	Pro
Pro	Glu 210		Leu	Val	Gly	Ser 215		Glu	Ser	: Lei	220		Trp	Ala	Tyr

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Gln Tyr Lys Ile Asn Gly Asn Gln Leu Leu Phe Ala Pro Asp Gly Asn 225 230 235 240

Trp Asn Glu Met Tyr Ser Gln Pro Leu Ser Cys Arg Tyr Leu Leu Lys
245 250 255

Ala Arg Ser Val Val Leu Pro Asp Gly Ser Arg Tyr Ser Val Asp Ile 260 265 270

Ile His Ser Lys Phe Ser His His Leu Leu Ser Phe Thr Pro Met Gly 275 280 285

Asn Leu Leu Thr Ser Asn Met Arg Cys Phe Ser Gly Phe Asp Ala Ile 290 295 300

Gly Ile Lys Asp Leu Glu Pro Leu Ser Arg Gly Met His Ser Cys Phe 305 310 315 320

Pro Val His His Asp Val Val Thr Lys Ile Tyr Leu Tyr Leu Arg Thr 325 330 335

Leu Lys Lys Pro Asp Lys Glu Ser Ala Glu Ala Lys Leu Arg Gln Leu 340 345 350

Ile Glu Lys Pro Thr Gly Arg Glu Ile Lys Phe Ile Glu Asp Phe Ser 355 360 365

Ser Leu Val Ile Asn Cys Gly Arg Ser Gly Ser Leu Leu Met Pro Asn 370 375 380

Ile Ser Lys Leu Val Ile Ser Phe Phe Cys Arg Met Met Pro Asn Ala 385 390 395 400

Leu Ala Arg Leu Ser Ser Ser Phe Arg Glu Cys Ser Leu Asp Ser Phe 405 410 415

Val Tyr Ser Leu Glu Pro Phe Asn Phe Ser Val Asn Leu Val Asp Ile 420 425 430

Thr Pro Asp Phe Phe Glu His Leu Phe Leu Phe Ser Cys Leu Asn Glu 435 440 445

Leu Ile Glu Glu Asp Val Glu Glu Val Met Asp Asn Ser Trp Phe Gly 450 455 460

Leu Gly Asp Leu Gln Phe Asn Arg Gln Arg Ala Pro Phe Phe Leu Gly 465 470 475 480

Ser Ser Tyr Trp Leu Asn Ser Lys Phe Ser Val Glu His Lys Phe Ser 485 490 495

Gly Thr Ile Asn Ser Gln Ile Met Gln Val Ile Leu Ser Leu Ile Pro 500 505 510

Phe Ser Asp Asp Pro Thr Phe Arg Pro Ser Ser Thr Glu Val Asn Leu 515 520 525

Ala Leu Ser Glu Val Lys Ala Ala Leu Glu Ala Thr Gly Gln Ser Lys 530 535 540

Leu Phe Arg Phe Leu Val Asp Asp Cys Ala Met Arg Glu Val Arg Ser 545 550 555 560

Ser Tyr Lys Val Gly Leu Phe Lys His Ile Lys Ala Leu Thr His Cys
565 570 575

Phe Asn Ser Cys Gly Leu Gln Trp Phe Leu Leu Arg Gln Arg Ser Asn 580 585 590

Leu Lys Phe Leu Lys Asp Arg Ala Ser Ser Phe Ala Asp Leu Asp Cys 595 600 605

Glu Val Ile Lys Val Tyr Gln Leu Val Thr Ser Gln Ala Ile Leu Pro 610 615 620

Glu Ala Leu Leu Ser Leu Thr Lys Val Phe Val Arg Asp Ser Asp Ser 625 630 635 640

Lys Gly Val Ser Ile Pro Arg Leu Val Ser Arg Asn Glu Leu Glu Glu 645 650 655

Leu Ala His Pro Ala Asn Ser Ala Leu Glu Glu Pro Gln Ser Val Asp
660 665 670

Cys Asn Ala Gly Arg Val Gln Ala Ser Val Ser Ser Ser Gln Gln Leu 675 680 685

Ala Asp Thr His Ser Leu Gly Ser Val Lys Ser Ser Ile Glu Thr Ala 690 695 700

Asn Lys Ala Phe Asn Leu Glu Glu Leu Arg Ile Met Ile Arg Val Leu 705 710 715 720

Pro Glu Asp Phe Asn Trp Val Ala Lys Asn Ile Gly Phe Lys Asp Arg
725 730 735

Leu Arg Gly Arg Gly Ala Ser Phe Phe Ser Lys Pro Gly Ile Ser Cys
740 745 750

His Ser Tyr Asn Gly Gly Ser His Thr Ser Leu Gly Trp Pro Lys Phe 755 760 765

Met Asp Gln Ile Leu Ser Ser Thr Gly Gly Arg Asn Tyr Tyr Asn Ser 770 780

Cys Leu Ala Gln Ile Tyr Glu Glu Asn Ser Lys Leu Ala Leu His Lys 785 790 795 800

Asp Asp Glu Ser Cys Tyr Glu Ile Gly His Lys Val Leu Thr Val Asn 805 810 815

Leu Ile Gly Ser Ala Thr Phe Thr Ile Ser Lys Ser Arg Asn Leu Val 820 825 830

Gly Gly Asn His Cys Ser Leu Thr Ile Gly Pro Asn Glu Phe Phe Glu 835 840 845

Met Pro Arg Gly Met Gln Cys Asn Tyr Phe His Gly Val Ser Asn Cys 850 860

- Thr Pro Gly Arg Val Ser Leu Thr Phe Arg Arg Gln Lys Leu Glu Asp 865 870 875 880
- Asp Asp Leu Ile Phe Ile Asn Pro Gln Val Pro Ile Glu Leu Asn His 885 890 895
- Glu Lys Leu Asp Arg Ser Met Trp Gln Met Gly Leu His Gly Ile Lys 900 905 910
- Lys Ser Ile Ser Met Asn Gly Thr Ser Phe Thr Ser Asp Leu Cys Ser 915 920 925
- Cys Phe Ser Cys His Asn Phe His Lys Phe Lys Asp Leu Ile Asn Asn 930 935 940
- Leu Arg Leu Ala Leu Gly Ala Gln Gly Leu Gly Gln Cys Asp Arg Val 945 950 955 960
- Val Phe Ala Thr Thr Gly Pro Gly Leu Ser Lys Val Leu Glu Met Pro 965 970 975
- Arg Ser Lys Lys Gln Ser Ile Leu Val Leu Glu Gly Ala Leu Ser Ile
 980 985 990
- Glu Thr Asp Tyr Gly Pro Lys Val Leu Gly Ser Phe Glu Val Phe Lys
- Gly Asp Phe His Ile Lys Lys Met Glu Glu Gly Ser Ile Phe Val Ile 1010 1015 1020
- Thr Tyr Lys Ala Pro Ile Arg Ser Thr Gly Arg Leu Arg Val His Ser 1025 1030 1035 1040
- Ser Glu Cys Ser Phe Ser Gly Ser Lys Glu Val Leu Leu Gly Cys Gln 1045 1050 1055
- Ile Glu Ala Cys Ala Asp Tyr Asp Ile Asp Asp Phe Asn Thr Phe Ser 1060 1065 1070
- Val Pro Gly Asp Gly Asn Cys Phe Trp His Ser Val Gly Phe Leu Leu 1075 1080 1085
- Ser Thr Asp Gly Leu Ala Leu Lys Ala Gly Ile Arg Ser Phe Val Glu 1090 1095 1100
- Ser Glu Arg Leu Val Ser Pro Asp Leu Ser Ala Pro Ala Ile Ser Lys 1105 1110 1115 1120
- Gln Leu Glu Glu Asn Ala Tyr Ala Glu Asn Glu Met Ile Ala Leu Phe 1125 1130 1135
- Cys Ile Arg His His Val Arg Pro Ile Val Ile Thr Pro Glu Tyr Glu 1140 1145 1150
- Val Ser Trp Lys Phe Gly Glu Gly Glu Trp Pro Leu Cys Gly Ile Leu 1155 1160 1165
- Cys Leu Lys Ser Asn His Phe Gln Pro Cys Ala Pro Leu Asn Gly Cys 1170 1175 1180

- Met Ile Thr Ala Ile Ala Ser Ala Leu Gly Arg Arg Glu Val Asp Val 1185 1190 1195 1200
- Leu Asn Tyr Leu Cys Arg Pro Ser Thr Asn His Ile Phe Glu Glu Leu
 1205 1210 1215
- Cys Gln Gly Gly Leu Asn Met Met Tyr Leu Ala Glu Ala Phe Glu 1220 1225 1230
- Ala Phe Asp Ile Cys Ala Lys Cys Asp Ile Asn Gly Glu Ile Glu Val 1235 1240 1245
- Ile Asn Pro Cys Gly Lys Ile Ser Ala Leu Phe Asp Ile Thr Asn Glu 1250 1255 1260
- His Ile Arg His Val Glu Lys Ile Gly Asn Gly Pro Gln Ser Ile Lys 1265 1270 1275 1280
- Val Asp Glu Leu Arg Lys Val Lys Arg Ser Ala Leu Asp Phe Leu Ser 1285 1290 1295
- Met Asn Gly Ser Lys Ile Thr Tyr Phe Pro Ser Phe Glu Arg Ala Glu 1300 1305 1310
- Lys Leu Gln Gly Cys Leu Leu Gly Gly Leu Thr Gly Val Ile Ser Asp 1315 1320 1325
- Glu Lys Phe Ser Asp Ala Lys Pro Trp Leu Ser Gly Ile Ser Thr Thr 1330 1335 1340
- Asp Ile Lys Pro Arg Glu Leu Thr Val Val Leu Gly Thr Phe Gly Ala 1345 1350 1355 1360
- Gly Lys Ser Phe Leu Tyr Lys Ser Phe Met Lys Arg Ser Glu Gly Lys 1365 1370 1375
- Phe Val Thr Phe Val Ser Pro Arg Arg Ala Leu Ala Asn Ser Ile Lys 1380 1385 1390
- Asn Asp Leu Glu Met Asp Asp Ser Cys Lys Val Ala Lys Ala Gly Arg 1395 1400 1405
- Ser Lys Lys Glu Gly Trp Asp Val Val Thr Phe Glu Val Phe Leu Arg 1410 1415 1420
- Lys Val Ala Gly Leu Lys Ala Gly His Cys Val Ile Phe Asp Glu Val 1425 1430 1435 1440
- Gln Leu Phe Pro Pro Gly Tyr Ile Asp Leu Cys Leu Leu Ile Ile Arg 1445 1450 1455
- Ser Asp Ala Phe Ile Ser Leu Ala Gly Asp Pro Cys Gln Ser Thr Tyr 1460 1465 1470
- Asp Ser Gln Lys Asp Arg Ala Ile Leu Gly Ala Glu Gln Ser Asp Ile 1475 1480 1485
- Leu Arg Leu Leu Glu Gly Lys Thr Tyr Arg Tyr Asn Ile Glu Ser Arg 1490 1495 1500

- Arg Phe Val Asn Pro Met Phe Glu Ser Arg Leu Pro Cys His Phe Lys 1505 1510 1515 1520
- Lys Gly Ser Met Thr Ala Ala Phe Ala Asp Tyr Ala Ile Phe His Asn 1525 1530 1535
- Met His Asp Phe Leu Leu Ala Arg Ser Lys Gly Pro Leu Asp Ala Val 1540 1545 1550
- Leu Val Ser Ser Phe Glu Glu Lys Lys Ile Val Gln Ser Tyr Phe Gly 1555 1560 1565
- Met Lys Gln Leu Thr Leu Thr Phe Gly Glu Ser Thr Gly Leu Asn Phe 1570 1580
- Lys Asn Gly Gly Ile Leu Ile Ser His Asp Ser Phe His Thr Asp Asp 1585 1590 1595 1600
- Arg Arg Trp Leu Thr Ala Leu Ser Arg Phe Ser His Asn Leu Asp Leu 1605 1610 1615
- Val Asn Ile Thr Gly Leu Arg Val Glu Ser Phe Leu Ser His Phe Ala 1620 1625 1630
- Gly Lys Pro Leu Tyr His Phe Leu Thr Ala Lys Ser Gly Glu Asn Val 1635 1640 1645
- Ile Arg Asp Leu Leu Pro Gly Glu Pro Asn Phe Phe Ser Gly Phe Asn 1650 1660
- Val Ser Ile Gly Lys Asn Glu Gly Val Arg Glu Glu Lys Leu Cys Gly 1665 1675 1680
- Asp Pro Trp Leu Lys Val Met Leu Phe Leu Gly Gln Asp Glu Asp Cys 1685 1690 1695
- Glu Val Glu Glu Met Glu Ser Glu Cys Ser Asn Glu Glu Trp Phe Lys 1700 1705 1710
- Thr His Ile Pro Leu Ser Asn Leu Glu Ser Thr Arg Ala Arg Trp Val 1715 1720 1725
- Gly Lys Met Ala Leu Lys Glu Tyr Arg Glu Val Arg Cys Gly Tyr Glu 1730 1740
- Met Thr Gln Gln Phe Phe Asp Glu His Arg Gly Gly Thr Gly Glu Gln 1745 1750 1755 1760
- Leu Ser Asn Ala Cys Glu Arg Phe Glu Ser Ile Tyr Pro Arg His Lys 1765 1770 1775
- Gly Asn Asp Ser Ile Thr Phe Leu Met Ala Val Arg Lys Arg Leu Lys 1780 1785 1790
- Phe Ser Lys Pro Gln Val Glu Ala Ala Lys Leu Arg Arg Ala Lys Pro 1795 1800 1805
- Tyr Gly Lys Phe Leu Leu Asp Ser Phe Leu Ser Lys Ile Pro Leu Lys 1810 1815 1820

- Ala Ser His Asn Ser Ile Met Phe His Glu Ala Val Gln Glu Phe Glu 1825 1830 1835 1840
- Ala Lys Lys Ala Ser Lys Ser Ala Ala Thr Ile Glu Asn His Ala Gly
 1845 1850 1855
- Arg Ser Cys Arg Asp Trp Leu Leu Asp Val Ala Leu Ile Phe Met Lys 1860 1865 1870
- Ser Gln His Cys Thr Lys Phe Asp Asn Arg Leu Arg Val Ala Lys Ala 1875 1880 1885
- Gly Gln Thr Leu Ala Cys Phe Gln His Ala Val Leu Val Arg Phe Ala 1890 1895 1900
- Pro Tyr Met Arg Tyr Ile Glu Lys Lys Leu Met Gln Ala Leu Lys Pro 1905 1910 1915 1920
- Asn Phe Tyr Ile His Ser Gly Lys Gly Leu Asp Glu Leu Asn Glu Trp 1925 1930 1935
- Val Arg Thr Arg Gly Phe Thr Gly Ile Cys Thr Glu Ser Asp Tyr Glu 1940 1945 1950
- Ala Phe Asp Ala Ser Gln Asp His Phe Ile Leu Ala Phe Glu Leu Gln 1955 1960 1965
- Ile Met Lys Phe Leu Gly Leu Pro Glu Asp Leu Ile Leu Asp Tyr Glu 1970 1975 1980
- Phe Ile Lys Ile His Leu Gly Ser Lys Leu Gly Ser Phe Ser Ile Met 1985 1990 1995 2000
- Arg Phe Thr Gly Glu Ala Ser Thr Phe Leu Phe Asn Thr Met Ala Asn 2005 2010 2015
- Met Leu Phe Thr Phe Leu Arg Tyr Glu Leu Thr Gly Ser Glu Ser Ile 2020 2025 2030
- Ala Phe Ala Gly Asp Asp Met Cys Ala Asn Arg Arg Leu Arg Leu Lys 2035 2040 2045
- Thr Glu His Glu Gly Phe Leu Asn Met Ile Cys Leu Lys Ala Lys Val 2050 2055 2060
- Gln Phe Val Ser Asn Pro Thr Phe Cys Gly Trp Cys Leu Phe Lys Glu 2065 2070 2075 2080
- Gly Ile Phe Lys Lys Pro Gln Leu Ile Trp Glu Arg Ile Cys Ile Ala 2085 2090 2095
- Arg Glu Met Gly Asn Leu Glu Asn Cys Ile Asp Asn Tyr Ala Ile Glu 2100 2105 2110
- Val Ser Tyr Ala Tyr Arg Leu Gly Glu Leu Ala Ile Glu Met Met Thr 2115 2120 2125
- Glu Glu Glu Val Glu Ala His Tyr Asn Cys Val Arg Phe Leu Val Arg 2130 2135 2140

Asn Lys His Lys Met Arg Cys Ser Ile Ser Gly Leu Phe Glu Ala Ile 2145 2150 2155 2160

Asp

The replicase of SEQ. ID. No. 3 has a molecular weight of about 240 to 246 kDa, preferably about 244 kDa.

Another DNA molecule of the present invention (RSPaV-1 ORF2) includes nucleotides 6578-7243 of SEQ. ID. No. 1. The DNA molecule of RSPaV-1 ORF2 encodes for a first protein or polypeptide of an RSPaV-1 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 4 as follows:

TAGTTAAAGC	ATTGTCAGCA	TTTGAGTTTG	TAGGTGTTTT	CAGTGTGCTT	60
TAGTCATTCA	TAGTGTGCCT	GGTAGTGGTA	AAAGTAGTTT	AATAAGGGAG	120
AGGATGAGAA	TTTCATAGCT	TTCACAGCAG	GTGTTCCAGA	CAGCCCTAAT	180
GGTACATTAA	GCCTTATTCT	CCAGGGTGTG	CAGTGCCAGG	GAAAGTTAAT	240
AGTACTTGTC	CGTCCAAGAT	TTTTCAGGTT	TTGATGTGCT	GTTCTCGGAC	300
ACATCAGCAT	TCCTAAAGAG	GCACATTTCA	TCAAGTCAAA	AACTTGTAGG	360
ATACTTGCAA	ATATCTTTCC	TCCTTCGGTT	TTAAGGTTAG	CAGTGACGGT	420
TCATTGTGGG	GTCGCCTTTT	ACACTAGATG	TTGAAGGGGT	GCTAATATGC	480
AGGCAGTGGA	TCTCGCTGTT	GCGCACAACT	CTGAATTCAA	ATTACCTTGT	540
GTTCAACTTT	TAACGTCGTA	ACTCTTTTGA	AATCAAGAGA	TCCAACCCCA	600
ACTGGTTTTA	CATTGCTGCT	ACAAGACACA	GGGAGAAATI	GATAATCATG	660
			•		663
	TAGTCATTCA AGGATGAGAA GGTACATTAA AGTACTTGTC ACATCAGCAT ATACTTGCAA TCATTGTGGG AGGCAGTGGA GTTCAACTTT	TAGTCATTCA TAGTGTGCCT AGGATGAGAA TTTCATAGCT GGTACATTAA GCCTTATTCT AGTACTTGTC CGTCCAAGAT ACATCAGCAT TCCTAAAGAG ATACTTGCAA ATATCTTTCC TCATTGTGGG GTCGCCTTTT AGGCAGTGGA TCTCGCTGTT GGTTCAACTTT TAACGTCGTA	TAGTCATTCA TAGTGTGCCT GGTAGTGGTA AGGATGAGAA TTTCATAGCT TTCACAGCAG GGTACATTAA GCCTTATTCT CCAGGGTGTG AGTACTTGTC CGTCCAAGAT TTTTCAGGTT ACATCAGCAT TCCTAAAGAG GCACATTTCA ATACTTGCAA ATATCTTTCC TCCTTCGGTT TCATTGTGGG GTCGCCTTTT ACACTAGATG AGGCAGTGGA TCTCGCTGTT GCGCACAACT GTTCAACTTT TAACGTCGTA ACTCTTTTGA	TAGTCATTCA TAGTGTGCT GGTAGTGGTA AAAGTAGTTT AGGATGAGAA TTTCATAGCT TTCACAGCAG GTGTTCCAGA GGTACATTAA GCCTTATTCT CCAGGGTGTG CAGTGCCAGG AGTACTTGTC CGTCCAAGAT TTTTCAGGTT TTGATGTGCT ACATCAGCAT TCCTAAAGAG GCACATTTCA TCAAGTCAAA ATACTTGCAA ATATCTTTCC TCCTTCGGTT TTAAGGTTAG TCATTGTGGG GTCGCCTTTT ACACTAGATG TTGAAGGGGT AGGCAGTGGA TCTCGCTGTT GCGCACAACT CTGAATTCAA GTTCAACTTT TAACGTCGTA ACTCTTTTGA AATCAAGAGA	TAGTTAAAGC ATTGTCAGCA TTTGAGTTTG TAGGTGTTTT CAGTGTGCTT TAGTCATTCA TAGTGTGCCT GGTAGTGGTA AAAGTAGTTT AATAAGGAG AGGATGAGAA TTTCATAGCT TTCACAGCAG GTGTTCCAGA CAGCCCTAAT GGTACATTAA GCCTTATTCT CCAGGGTGTG CAGTGCCAGG GAAAGTTAAT AGTACTTGTC CGTCCAAGAT TTTTCAGGTT TTGATGTGCT GTTCTCGGAC ACATCAGCAT TCCTAAAGAG GCACATTTCA TCAAGTCAAA AACTTGTAGG ATACTTGCAA ATATCTTTCC TCCTTCGGTT TTAAGGTTAG CAGTGACGGT TCATTGTGGG GTCGCCTTTT ACACTAGATG TTGAAGGGGT GCTAATATGC AGGCAGTGGA TCTCGCTGTT GCGCACAACT CTGAATTCAA ATTACCTTGT GTTCAACTTT TAACGTCGTA ACTCTTTTGA AATCAAGAGA TCCAACCCCA ACTGGTTTTA CATTGCTGCT ACAAGACACA GGGAGAAATT GATAATCATG

The first protein or polypeptide of the RSPaV-1 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 5 as follows:

Met Asn Asn Leu Val Lys Ala Leu Ser Ala Phe Glu Phe Val Gly Val 15

Phe Ser Val Leu Lys Phe Pro Val Val Ile His Ser Val Pro Gly Ser 25

Gly Lys Ser Ser Leu Ile Arg Glu Leu Ile Ser Glu Asp Glu Asn Phe 45

Ile Ala Phe Thr Ala Gly Val Pro Asp Ser Pro Asn Leu Thr Gly Arg 60

Tyr 65	Ile	Lys	Pro	Tyr	Ser 70	Pro	Gly	Cys	Ala	Val 75	Pro	Gly	Lys	Val	Asn 80
Ile	Leu	Asp	Glu	Tyr 85	Leu	Ser	Val	Gln	Asp 90	Pḥe	Ser	Gly	Phe	Asp 95	Val
Leu	Phe	Ser	Asp 100	Pro	Tyr	Gln	Asn	Ile 105	Ser	Ile	Pro	Lys	Glu 110	Ala	His
Phe	Ile	Lys 115	Ser	Lys	Thr	Cys	Arg 120	Phe	Gly	Val	Asn	Thr 125	Cys	Lys	Tyr
Leu	Ser 130	Ser	Phe	Gly	Phe	Lys 135	Val	Ser	Ser	Asp	Gly 140	Leu	Asp	Lys	Val
Ile 145	Val	Gly	Ser	Pro	Phe 150		Leu	Asp	Val	Glu 155	Gly	Val	Leu	Ile	Cys 160
Phe	Gly	Lys	Glu	Ala 165		Asp	Leu	Ala	Val	Ala	His	Asn	Ser	Gl u 175	Phe
Lys	Leu	Pro	Cys 180		Val	Arg	Gly	Ser 185	Thr	Phe	Asn	Val	. Val 190	Thi	: Leu
Leu	Lys	Ser 195		Asp	Pro	Thr	200		a Asp	Arc	His	205	Phe	ту	r Ile
Ala	Ala 210		Arç	, His	arç	Glu 215		Let	ı Ile	e Ile	221	Gli O	n .		

The first protein or polypeptide of the RSPaV-1 triple gene block has a molecular weight of about 20 to 26 kDa, preferably 24.4 kDa.

Another DNA molecule of the present invention (RSPaV-1 ORF3) includes nucleotides 7245-7598 of SEQ. ID. No. 1. The DNA molecule of RSPaV-1 ORF3 encodes for a second protein or polypeptide of the triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 6 as follows:

ATGCCTTTTC	AGCAGCCTGC	GAATTGGGCA	AAAACCATAA	CTCCATTGAC	AGTTGGCTTG	60
GGCATTGGGC	TTGTGCTGCA	TTTTCTGAGG	AAGTCAAATC	TACCTTATTC	AGGGGACAAC	120
ATCCATCAAT	TCCCTCACGG	TGGGCGTTAC	AGGGACGGTA	CAAAAAGTAT	AACTTACTGT	180
GGTCCAAAGC	AATCCTTCCC	CAGCTCTGGG	ATATTCGGCC	AATCTGAGAA	TTTTGTGCCC	24
TTAATGCTTG	TCATAGGTCT	AATCGCATTC	ATACATGTAT	TGTCTGTTTG	GAATTCTGGT	. 30
CTTGGTAGGA	ATTGTAATTG	CCATCCAAAT	CCTTGCTCAT	GTAGACAGCA	G	35

The second protein or polypeptide of the RSPaV-1 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 7 as follows:

Met 1	Pro	Phe	Gln	Gln 5	Pro	Ala	Asn	Trp	Ala 10	Lys	Thr	Ile	Thr	Pro 15	Leu
Thr	Val	Gly	Leu 20	Gly	Ile	Gly	Leu	Val 25	Leu	His	Phe	Leu	Arg 30	Lys	Ser
Asn	Leu	Pro 35	Tyr	Ser	Gly	Asp	Asn 40	Ile	His	Gln	Phe	Pro 45	His	Gly	Gly
Arg	Tyr 50	Arg	Asp	Gly	Thr	Lys 55	Ser	Ile	Thr	Tyr	Cys 60	Gly	Pro	Lys	Gln
Ser 65	Phe	Pro	Ser	Ser	Gly 70	Ile	Phe	Gly	Gln	Ser 75	Glu	Asn	Phe	Val	Pro 80
Leu	Met	Leu	Val	Ile 85	Gly	Leu	Ile	Ala	Phe 90	Ile	His	Val	Leu	Ser 95	Val
Trp	Asn	Ser	Gly 100		GJA	Arg	Asn	Cys 105		Cys	His	Pro	Asn 110		Cys
Ser	Cys	Arg 115		Gln						•					

The second protein or polypeptide of the RSPaV-1 triple gene block has a molecular weight of about 10 to 15 kDa, preferably 12.8 kDa.

Yet another DNA molecule of the present invention (RSPaV-1 ORF4) includes nucleotides 7519-7761 of SEQ. ID. No. 1. The DNA molecule of RSPaV-1 ORF4 encodes for a third protein or polypeptide of the RSPaV-1 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 8 as follows:

ATGTATTGTC	TGTTTGGAAT	TCTGGTCTTG	GTAGGAATTG	TAATTGCCAT	CCAAATCCTT	60
GCTCATGTAG	ACAGCAGTAG	TGGCAACCAC	CAAGGTTGCT	TCATTAGGGC	CACTGGAGAG	120
TCAATTTTGA	TTGAAAACTG	CGGCCCAAGT	GAGGCCCTTG	CATCCACTGT	GAAGGAGGTG	180
CTGGGAGGTT	TGAAGGCTTT	AGGGGTTAGC	CGTGCTGTTG	AAGAAATTGA	TTATCATTGT	240

The third protein or polypeptide of the RSPaV-1 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 9 as follows:

	Tyr					Ile	Leu				Gly			Ile 15	Ala
Ile	Gln	Ile	Leu 20		His	Val					Gly			Gĺn	Gly
Cys		Ile 35	Arg	Ala	Thr		Glu 40	Ser	Ile	Leu	Ile	Glu 45	Asn	Cys	Gly

Pro Ser Glu Ala Leu Ala Ser Thr Val Lys Glu Val Leu Gly Gly Leu 50 55 60

Lys Ala Leu Gly Val Ser Arg Ala Val Glu Glu Ile Asp Tyr His Cys 65 70 75 80

The third protein or polypeptide of the RSPaV-1 triple gene block has a molecular weight of about 5 to 10 kDa, preferably 8.4 kDa.

Still another DNA molecule of the present invention (RSPaV-1 ORF5) includes nucleotides 7771-8550 of SEQ. ID. No. 1. The DNA molecule of RSPaV-1 ORF5 encodes for a RSPaV-1 coat protein and comprises a nucleotide sequence corresponding to SEQ. ID. No. 10 as follows:

ATGGCAAGTC	AAATTGGGAA	ACTCCCCGGT	GAATCAAATG	AGGCTTTTGA	AGCCCGGCTA	60
AAATCGCTGG	AGTTAGCTAG	AGCTCAAAAG	CAGCCGGAAG	GTTCTAATGC	ACCACCTACT	120
CTCAGTGGCA	TTCTTGCCAA	ACGCAAGAGG	ATTATAGAGA	ATGCACTTTC	AAAGACGGTG	180
GACATGAGGG	AGGTTTTGAA	ACACGAAACG	GTGGTGATTT	CCCCAAATGT	CATGGATGAA	240
GGTGCAATAG	ACGAGCTGAT	TCGTGCATTT	GGTGAATCTG	GCATAGCTGA	AAGCGTGCAA	300
TTTGATGTGG	CCATAGATAT	AGCACGTCAC	TGCTCTGATG	TTGGTAGCTC	CCAGAGTTCA	360
ACCCTGATTG	GCAAGAGTCC	ATTTTGTGAC	CTAAACAGAT	CAGAAATAGC	TGGGATTATA	420
AGGGAGGTGA	CCACATTACG	TAGATTTTGC	ATGTACTATG	CAAAAATCGT	GTGGAACATC	480
CATCTGGAGA	CGGGGATACC	ACCAGCTAAC	TGGGCCAAGA	AAGGATTTAA	TGAGAATGAA	540
AAGTTTGCAG	CCTTTGATTT	TTTCTTGGGA	GTCACAGATG	AGAGTGCGCT	TGAACCAAAG	600
GGTGGAATTA	AAAGAGCTCC	AACGAAAGCT	GAGATGGTTG	CTAATATCGC	CTCTTTTGAG	660
GTTCAAGTGC	TCAGACAAGC	TATGGCTGAA	GGCAAGCGGA	GTTCCAACCI	TGGAGAGATT	720
AGTGGTGGAA	CGGCTGGTGC	ACTCATCAAC	AACCCCTTT	CAAATGTTAC	ACATGAA	777

The RSPaV-1 coat protein has a deduced amino acid sequence corresponding to SEQ. ID. No. 11 as follows:

Lys	Arg 50	Ile	Ile	Glu		Ala 55	Leu :	Ser	Lys	Thr	Val .	Asp	Met .	Arg	Glu
Val 65	Leu	Lys	His	Glu	Thr 70	Val	Val	Ile	Ser	Pro 75	Asn	Val	Met	Asp	Glu 80
Gly	Ala	Ile	Asp	Glu 85	Leu	Ile	Arg	Ala	Phe 90	Gly	Glu	Ser	Gly	11e 95	Ala
Glu	Ser	Val	Gln 100	Phe	Asp	Val	Ala	Ile 105	Asp	Ile	Ala	Arg	His 110	Cys	Ser
Asp	Val	Gly 115	Ser	Ser	Gln	Ser	Ser 120	Thr	Leu	Ile	Gly	Lys 125	Ser	Pro	Phe
Cys	Asp 130		Asn	Arg	Ser	Glu 135	Ile	Ala	Gly	Ile	11e 140	Arg	Glu	Val	Thr
Thr 145	Leu	Arg	Arg	Phe	Cys 150	Met	Tyr	Tyr	Ala	Lys 155	Ile	Val	Trp	Asn	Ile 160
His	Leu	Glu	Thr	Gly 165	_	Pro	Pro	Ala	Asn 170	Trp	Ala	Lys	Lys	Gly 175	Phe
Asn	Glu	Asn	Glu 180		Phe	Ala	Ala	Phe 185	Asp	Phe	. Phe	Leu	Gly 190	Val	Thr
Asp	Glu	Ser 195		Leu	Glu	Pro	Lys 200	Gly	Gly	, Ile	e Lys	Arg 205	Ala	Pro	Thr
Lys	Ala 210		Met	. Val	Ala	Asn 215		Ala	a Ser	Phe	e Glu 220	ī Val	Glr	va.	L Leu
Arg 225		n Ala	a Met	Ala	230		/ Lys	Ar	g Sei	23	r Asr 5	ı Lei	ı Gʻl?	/ Gl	11e 240
Ser	c Gly	y Gl	y Thi	r Ala 24		/ Ala	a Leu	ı Ile	e Ası 25	n As O	n Pro	o Phe	e Se	25	n Val
Th	r His	s Gl	u									i			

IIII III3 OIG

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The RSPaV-1 coat protein has a molecular weight of about 25 to 30 kDa, preferably 28 kDa.

The DNA molecule which constitutes the substantial portion of the RSPaV strain RSP47-4 genome comprises the nucleotide sequence corresponding to SEQ. ID. No. 12 as follows:

GGCTGGGCAA ACTTTGG	CCT GCTTTCAACA	CGCCGTCTTG	GTTCGCTTTG	CACCCTACAT	60
GCGATACATT GAAAAGA	AGC TTGTGCAGGC	ATTGAAACCA	AATTTCTACA	TTCATTCTGG	120
CAAAGGTCTT GATGAGC	TAA GTGAATGGGT	TAGAGCCAGA	GGTTTCACAG	GTGTGTGTAC	180
TCAGTCAGAC TATGAAG	CTT TTGATGCATO	CCAAGATCAT	TTCATCCTGG	CATTTGAACT	. 240

SCAAATCATG AGATTTTTAG GACTGCCAGA AGATCTGATT TTAGATTAT	G AGTTCATCAA 300
AATTCATCTT GGGTCAAAGC TTGGCTCTTT TGCAATTATG AGATTCACA	G GTGAGGCAAG 360
CACCTTCCTA TTCAATACTA TGGCCAACAT GCTATTCACT TTCCTGAGG	T ATGAGTTGAC 420
AGGTTCTGAA TCAATTGCAT TTGCTGGAGA TGATATGTGT GCTAATCGC	A GGTTAAGACT 480
CAAGACTGAG CACGCCGGCT TTCTAAACAT GATCTGTCTC AAAGCTAAG	G TGCAGTTTGT 540
CACAAATCCC ACCTTCTGTG GATGGTGTTT GTTTAAAGAG GGAATCTTT	A AAAAACCCCA 600
CTCATTTGG GAAAGGATCT GCATTGCTAG GGAAATGGGT AACTTGGAC	A ATTGCATTGA 660
CAATTACGCA ATTGAGGTGT CTTATGCTTA CAGACTTGGG GAATTGTCC	A TAGGCGTGAT 720
GACTGAGGAG GAAGTTGAAG CACATTCTAA CTGCGTGCGT TTCCTGGTT	C GCAATAAGCA 780
CAAGATGAGG TGCTCAATTT CTGGTTTGTT TGAAGTAATT GTTTAGGCC	T TAAGTGTTTG 840
GCATGGTGTG AGTATTATGA ATAACTTAGT CAAAGCTTTG TCTGCTTTT	G AATTTGTTGG 900
TGTGTTTTGT GTACTTAAAT TTCCAGTTGT TGTTCACAGT GTTCCAGGT	'A GCGGTAAAAG 960
TAGCCTAATA AGGGAGCTCA TTTCTGAAGA CGAGGCTTTT GTGGCCTTI	CA CAGCAGGTGT 1020
GCCAGACAGT CCAAATCTGA CAGGGAGGTA CATCAAGCCC TACGCTCCA	AG GGTGTGCAGT 1080
GCAAGGGAAA ATAAACATAC TTGATGAGTA CTTGTCTGTC TCTGATAC	TT CTGGCTTTGA 1140
TGTGCTGTTC TCAGACCCTT ACCAGAATGT CAGCATTCCA AGGGAGGC	AC ACTTCATAAA 1200
AACCAAAACC TGTAGGTTTG GTACCAACAC CTGCAAGTAC CTTCAATC	TT TTGGCTTTAA 1260
TGTTTGTAGT GATGGGGTGG ATAAAGTTGT TGTAGGGTCG CCATTTGA	AC TGGAGGTTGA 1320
GGGGGTTCTC ATTTGCTTTG GAAAGGAGGC TGTAGATCTA GCAGTTGC	AC ACAATTCTGA 1380
CTTCAAGTTG CCCTGCGAGG TGCGGGGTTC AACATTTGAC GTTGTAAC	GT TATTGAAGTC 1440
CAGGGATCCA ACTTCAGAAG ATAAGCATTG GTTCTACGTT GCAGCCAC	AA GGCATCGAAG 1500
TAAACTGATA ATAATGCAGT AAAATGCCTT TTCAGCAACC TGCCAACT	GG GCTAAGACCA 1560
TAACTCCATT AACTATTGGT TTGGGCATTG GGTTGGTTCT GCACTTCT	TA AGGAAATCAA 1620
ATCTGCCATA TTCAGGAGAC AATATTCACC AGTTCCCACA CGGAGGGC	AT TACAGGGACG 1680
GCACGAAGAG TATAACCTAT TGTGGCCCTA GGCAGTCATT CCCAAGCT	CA GGAATATTCG 1740
GTCAGTCTGA AAATTTCGTA CCTCTAATAT TGGTCGTGAC TCTGGTCG	SCT TTTATACATG 1800
CGTTATCTCT TTGGAATTCT GGTCCTAGTA GGAGTTGCAA TTGCCATC	CCA AATCCTTGCA 1860
CATGTAGACA GCAGTAGTGG CAACCATCAA GGCTGTTTCA TAAGAGCC	CAC CGGGGAGTCA 1920
ATAGTAATTG AGAATTGTGG GCCGAGCGAG GCCCTAGCTG CTACAGTG	CAA AGAGGTGTTG 198
GGCGGTCTAA AGGCTTTAGG GGTTAGCCAA AAGGTTGATG AAATTAA'	TTA CAGTTGTTGA 204

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G	ACAGTTGAA	TGGCAAGTCA	AGTTGGAAAA	TTGCCTGGCG	AATCAAATGA	AGCATATGAG	2100
G	CTAGACTCA	AGGCTTTAGA	GTTAGCAAGG	GCCCAAAAAG	CTCCAGAAGT	CTCCAACCAA	2160
c	CTCCCACAC	TTGGAGGCAT	TCTAGCCAAA	AGGAAAAGAG	TGATTGAGAA	TGCACTCTCA	2220
P	AGACAGTGG	ATATGCGTGA	AGTCTTAAGG	CATGAATCTG	TTGTACTCTC	CCCGAATGTA	2280
7	ATGGACGAGG	GAGCAATAGA	CGAGCTGATT	CGTGCCTTTG	GGGAGTCGGG	CATAGCTGAA	2340
7	AATGTGCAGT	TTGATGTTGC	AATAGACATT	GCTCGCCACT	GTTCTGATGT	GGGGAGCTCT	2400
(CAGAGGTCAA	CCCTTATTGG	TAAAAGCCCC	TTCTGTGAGT	TAAATAGGTC	TGAAATTGCC	2460
(GGAATAATAA	GGGAGGTGAC	CACGCTGCGC	AGATTTTGCA	TGTACTACGC	AAAGATTGTG	2520
•	TGGAACATCC	ATTTGGAGAC	GGGAATACCA	CCAGCTAATT	GGGCCAAGAA	AGGATTTAAT	2580
•	GAGAATGAAA	AGTTTGCAGC	CTTTGACTTC	TTCCTTGGAG	TCACAGATGA	AAGCGCGCTT	2640
	GAGCCTAAGG	GTGGAGTCAA	GAGAGCTCCA	ACAAAAGCAG	;		2680

The RSP47-4 strain contains five open reading frames (i.e., ORF1-5). ORF1 and ORF5 are only partially sequenced. RSP47-4 is 79% identical in nucleotide sequence to the corresponding region of RSPaV-1. The amino acid sequence identities between the corresponding ORFs of RSP47-4 and RSPaV-1 are: 94.1% for ORF1, 88.2% for ORF2, 88.9% for ORF3, 86.2% for ORF4, and 92.9% for ORF5. The nucleotide sequences of the five potential ORFs of RSP47-4 are given below.

Another DNA molecule of the present invention (RSP47-4 incomplete ORF1) includes nucleotides 1-768 of SEQ. ID. No. 12. This DNA molecule is believed to code for a polypeptide portion of a RSP47-4 replicase and comprises a nucleotide sequence corresponding to SEQ. ID. No. 13 as follows:

ATGCGATACA TTGAAAAGAA GCTTGTGCAG GCATTGAAAC CAAATTTCTA CATTCATTCT 60 GGCAAAGGTC TTGATGAGCT AAGTGAATGG GTTAGAGCCA GAGGTTTCAC AGGTGTGTGT 120 ACTGAGTCAG ACTATGAAGC TTTTGATGCA TCCCAAGATC ATTTCATCCT GGCATTTGAA 180 CTGCAAATCA TGAGATTTTT AGGACTGCCA GAAGATCTGA TTTTAGATTA TGAGTTCATC 240 AAAATTCATC TTGGGTCAAA GCTTGGCTCT TTTGCAATTA TGAGATTCAC AGGTGAGGCA 300 AGCACCTTCC TATTCAATAC TATGGCCAAC ATGCTATTCA CTTTCCTGAG GTATGAGTTG 360 ACAGGTTCTG AATCAATTGC ATTTGCTGGA GATGATATGT GTGCTAATCG CAGGTTAAGA 420 CTCAAGACTG AGCACGCCGG CTTTCTAAAC ATGATCTGTC TCAAAGCTAA GGTGCAGTTT 480 GTCACAAATC CCACCTTCTG TGGATGGTGT TTGTTTAAAG AGGGAATCTT TAAAAAACCC 540

CAGCT	CATT	T GG	GAAA	GGAT	CTG	TTA	CT F	AGGG?)TAA!	G GI	AACT	TGGA	CA	ATTGO	CATT		600
GACAA	TTAC	G CA	ATTG.	AGGT	GTC	TATO	CT 1	CACAC	GACTI	rg gg	GAAT	TGTC	CA	PAGG	CGTG		660
ATGAC	TGAG	G AG	GAAG	TTGA	AGC	ACATI	CT A	AACTO	CGT	GC GT	TTCC	TGGT	TC	GCAA'	TAA G		720
CACAA	GATG	A GG	TGCT	CAAT	TTC'	rggt:	rtg :	rttg/	AAGT?	AA TI	rgttt	'A		-			767
The p	olype	ptide	has	a ded	uced	amin	o aci	d seq	uenc	е соп	espoi	nding	to S	EQ.	ID. N	lo.	
14 as	follo	ws:				•					:						
•	Met 1	Arg	Tyr	Ile	Glu 5	Lys :	Lys	Leu		Gln 1 10	Ala I	Leu I	ys		Asn 15	Phe	
	Tyr	Ile	His	Ser 20	Gly	Lys	Gly		Asp 25	Glu :	Leu S	Ser (Slu	Trp 30	Val	Arg	-
· .:	Ala	Arg	Gly 35	Phe	Thr	Gly	Val	Cys 40	Thr	Glu	Ser i		Tyr 45	Glu	Ala	Phe	•
	Asp	Ala 50	Ser	Gln	Asp	His	Phe 55	Ile	Leu	Ala	Phe	Glu i 60	Leu	Gln	Ile	Met	•
	Arg 65	Phe	Leu	Gly	Leu	Pro 70	Glu	Asp	Leu	Ile	Leu 75	Asp '	Tyr	Glu	Phe	Ile 80	
-	Lys	Ile	His	Leu	Gly 85	Ser	Lys	Leu	Gly	Ser 90	Phe	Ala	Ile	Met	Arg 95	Phe	
	Thr	Gly	Glu	Ala 100	Ser	Thr	Phe	Leu	Phe 105	Asn	Thr	Met	Ala	Asn 110	Met	Leu	
	Phe	Thr	Phe 115	Leu	Arg	Tyr	Glu	Leu 120	Thr	Gly:	Ser	Glu:	Ser 125	Ile	Ala	Phe	
	Ala	Gly 130	_	Asp	Met	Cys	Ala 135	Asn	Arg	Arg	Leu	Arg 140	Leu	Lys	Thr	Glu	- -
	His 145		Gly	Phe	Leu	Asn 150	Met	Ile	Cys	Leu	Lys 155	Ala	Lys	Val	Gln	Phe 160	
·	Val	Thr	Asn	Pro	Thr 165		Cys	Gly	Trp	Cys 170		Phe	Lys	Glu	Gly 175	lle	
	Phe	Lys	Lys	Pro 180		Leu	Ile	Trp	Glu 185		Ile	Cys	Ile	Ala 190	Arç	g Glu	1
·	Met	Gly	Asn 195		Asp	Asn	Cys	11e		Asn	Tyr	Ala	11e 205		ı Val	l Ser	<u>.</u>
	Туг	Ala 210		Arg	, Leu	Gly	Glu 215		ser	Ile	Gly	Val 220		Thi	c Gl	u Glu	נ
	Glu 225		l Glu	ı Ala	a His	Ser 230		ı Cys	s Val	l Arg	Phe 235		Va:	l Ar	g As	n Ly:	s 0
	His	E Lys	s Met	. Arç	Cys 245		: Ile	e Sei	r Gly	y Let 250	ı Phe	Glu	ı Va	111	e Va 25	1 5	

Another DNA molecule of the present invention (RSP47-4 ORF2) includes nucleotides 857-1522 of SEQ. ID. No. 12. This DNA molecule codes for a first protein or polypeptide of an RSP47-4 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 15 as follows:

ATGAATAACT	TAGTCAAAGC	TTTGTCTGCT	TTTGAATTTG	TTGGTGTGTT	TTGTGTACTT	60
AAATTTCCAG	TTGTTGTTCA	CAGTGTTCCA	GGTAGCGGTA	AAAGTAGCCT	AATAAGGGAG	120
CTCATTTCTG	AAGACGAGGC	TTTTGTGGCC	TTTACAGCAG	GTGTGCCAGA	CAGTCCAAAT	180
CTGACAGGGA	GGTACATCAA	GCCCTACGCT	CCAGGGTGTG	CAGTGCAAGG	GAAAATAAAC	240
ATACTTGATG	AGTACTTGTC	TGTCTCTGAT	ACTTCTGGCT	TTGATGTGCT	GTTCTCAGAC	300
CCTTACCAGA	ATGTCAGCAT	TCCAAGGGAG	GCACACTTCA	TAAAAACCAA	AACCTGTAGG	360
					TAGTGATGGG	420
					TCTCATTTGC	480
TTTGGAAAGG	AGGCTGTAGA	TCTAGCAGTT	GCACACAATT	CTGACTTCAA	GTTGCCCTGC	540
				• •	TCCAACTTCA	600
- T					GATAATAATG	660
CAGTAA					er en	666

The first protein or polypeptide of the RSP47-4 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 16 as follows:

Met Asn Asn Leu Val Lys Ala Leu Ser Ala Phe Glu Phe Val Gly Val 15

Phe Cys Val Leu Lys Phe Pro Val Val Val His Ser Val Pro Gly Ser 30

Gly Lys Ser Ser Leu Ile Arg Glu Leu Ile Ser Glu Asp Glu Ala Phe 45

Val Ala Phe Thr Ala Gly Val Pro Asp Ser Pro Asn Leu Thr Gly Arg 50

Tyr Ile Lys Pro Tyr Ala Pro Gly Cys Ala Val Gln Gly Lys Ile Asn 80

Ile Leu Asp Glu Tyr Leu Ser Val Ser Asp Thr Ser Gly Phe Asp Val 95

Leu Phe Ser Asp Pro Tyr Gln Asn Val Ser Ile Pro Arg Glu Ala His

Phe	Ile	Lys 115	Thr	Lys	Thr	Cys	Arg 120	Phe	Gly	Thr	Asn	Thr 125		Lys	Tyr
Leu	Gln 130	Ser	Phe	Gly	Phe	Asn 135	Val	Cys	Ser	Asp	Gly 140	Val	Asp	Lys	Val
Val 145	Val	Gly	Ser	Pro	Phe 150	Glu	Leu	Glu	Val	Glu 155	Gly	Val	Leu	Ile	Cys 160
Phe	Gly	Lys	Glu	Ala 165	Val	Asp	Leu	Ala	Val 170	Ala	His	Asn	Ser	Asp 175	Phe
Lys	Leu	Pro	Cys 180	Glu	Val	Arg	Gly	Ser 185	Thr	Phe	Asp	Val	Val 190	Thr	Leu
Leu	Lys	Ser 195	Arg	Asp	Pro	Thr	Ser 200	Glu	Asp	Lys	His	Trp 205	Phe	Tyr	Va]
Ala	Ala 210	Thr	Arg	His	Arg	Ser 215	Lys	Leu	Ile	Ile	Met 220				

The first protein or polypeptide of the RSP47-4 triple gene block has a molecular weight of about 20 to 26 kDa., preferably 24.3 kDa.

Another DNA molecule of the present invention (RSP47-4 ORF3) includes nucleotides 1524-1877 of SEQ. ID. No. 12. This DNA molecule codes for a second protein or polypeptide of the RSP47-4 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 17 as follows:

ATGCCTTTTC	AGCAACCTGC	CAACTGGGCT	AAGACCATAA	CTCCATTAAC	TATTGGTTTG	60
GGCATTGGGT	TGGTTCTGCA	CTTCTTAAGG	AAATCAAATC	TGCCATATTC	AGGAGACAAT	120
ATTCACCAGT	TCCCACACGG	AGGGCATTAC	AGGGACGGCA	CGAAGAGTAT	AACCTATTGT	180
GGCCCTAGGC	AGTCATTCCC	AAGCTCAGGA	ATATTCGGTC	AGTCTGAAAA	TTTCGTACCT	240
CTAATATTGG	TCGTGACTCT	GGTCGCTTTT	ATACATGCGT	TATCTCTTTG	GAATTCTGGT	300
CCTAGTAGGA	GTTGCAATTG	CCATCCAAAT	CCTTGCACAT	GTAGACAGCA	GTAG	354

The second protein or polypeptide of the RSP47-4 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 18 as follows:

Met Pro Phe Gln Gln Pro Ala Asn Trp Ala Lys Thr Ile Thr Pro Leu 15

Thr Ile Gly Leu Gly Ile Gly Leu Val Leu His Phe Leu Arg Lys Ser 20

Asn Leu Pro Tyr Ser Gly Asp Asn Ile His Gln Phe Pro His Gly Gly 45

His	Tyr 50	Arg	Asp	Gly	Thr	Lys 55	Ser	Ile	Thr	Tyr	Cys 60	Gly	Pro	Arg	Gln
Ser 65	Phe	Pro	Ser	Ser	Gly 70	Île	Phe	Gly	Gln	Ser 75	Glu	Asn	Phe	Val	Pro 80
Leu	Ile	Leu	Val	Val 85	Thr	Leu	Val	Ala	Phe 90	Ile	His	Ala	Leu	Ser 95	Leu
Trp	Asn	Ser	Gly 100	Pro	Ser	Arg	Ser	Cys 105	Asn	Cys	His	Pro	Asn 110	Pro	Cys
Thr		Arg	Gln	Gĺn						•					

The second protein or polypeptide of the RSP47-4 triple gene block has a molecular weight of about 10 to 15 kDa., preferably 12.9 kDa.

Another DNA molecule of the present invention (RSP47-4 ORF4) includes nucleotides 1798-2040 of SEQ. ID. No. 12. This DNA molecule codes for a third protein or polypeptide of the RSP47-4 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 19 as follows:

ATGCGTTATC	TCTTTGGAAT	TCTGGTCCTA	GTAGGAGTTG	CAATTGCCAT	CCAAATCCTT	60
GCACATGTAG	ACAGCAGTAG	TGGCAACCAT	CAAGGCTGTT	TCATAAGAGC	CACCGGGGAG	120
TCAATAGTAA	TTGAGAATTG	TGGGCCGAGC	GAGGCCCTAG	CTGCTACAGT	CAAAGAGGTG	180
TTGGGCGGTC	TAAAGGCTTT	AGGGGTTAGC	CAAAAGGTTG	ATGAAATTAA	TTACAGTTGT	240
TGA	•					243

The third protein or polypeptide of the RSP47-4 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 20 as follows:

Met 1	Arg	Tyr	Leu	Phe 5	Gly	Ile	Leu	Val	Leu 10	Val	Gly	Val	Ala	Ile 15	Ala
Ile	Gln	Ile	Leu 20	Ala	His	Val	Asp	Ser 25	Ser	Ser	Gly	Asn	His 30	Gln	GӀ҈у
Cys	Phe	Ile 35	Arg	Ala	Thr	Gly	Glu 40	Ser	Ile	Val	Ile	Glu 45	Asn	Cys	Gly
Pro	Ser 50	Glu	Ala	Leu	Ala	Ala 55	Thr	Val	Lys	Glu	Val 60	Leu	Gly	Gly	Leu
Lys 65	Ala	Leu	Gly	Val	Ser 70	Gln	Lys	Val	Asp	Glu 75	Ile	Asn	Tyr	Ser	Cys 80

The third protein or polypeptide of the RSP47-4 triple gene block has a molecular weight of about 5 to 10 kDa., preferably 8.3 kDa.

Yet another DNA molecule of the present invention (RSP47-4 ORF5) includes nucleotides 2050-2680 of SEQ. ID. No. 12. This DNA molecule codes for a partial RSP47-4 coat protein or polypeptide and comprises a nucleotide sequence corresponding to SEQ. ID. No. 21 as follows:

ATGGCAAGTC AAGTTGGAAA ATTGCCTGGC GAATCAAATG AAGCATATGA GGCTAGACTC 60 AAGGCTTTAG AGTTAGCAAG GGCCCAAAAA GCTCCAGAAG TCTCCAACCA ACCTCCCACA 120 CTTGGAGGCA TTCTAGCCAA AAGGAAAAGA GTGATTGAGA ATGCACTCTC AAAGACAGTG 180 GATATGCGTG AAGTCTTAAG GCATGAATCT GTTGTACTCT CCCCGAATGT AATGGACGAG 240 GGAGCAATAG ACGAGCTGAT TCGTGCCTTT GGGGAGTCGG GCATAGCTGA AAATGTGCAG 300 TTTGATGTTG CAATAGACAT TGCTCGCCAC TGTTCTGATG TGGGGAGCTC TCAGAGGTCA 360 ACCCTTATTG GTAAAAGCCC CTTCTGTGAG TTAAATAGGT CTGAAATTGC CGGAATAATA 420 AGGGAGGTGA CCACGCTGCG CAGATTTTGC ATGTACTACG CAAAGATTGT GTGGAACATC 480 CATTTGGAGA CGGGAATACC ACCAGCTAAT TGGGCCAAGA AAGGATTTAA TGAGAATGAA 540 AAGTTTGCAG CCTTTGACTT CTTCCTTGGA GTCACAGATG AAAGCGCGCT TGAGCCTAAG 600 GGTGGAGTCA AGAGAGCTCC AACAAAGCA G 631

The polypeptide has a deduced amino acid sequence corresponding to SEQ. ID. No. 22 as follows:

Asp	Val	Gly 115	Ser	Ser	Gln	Arg	Ser 120	Thr	Leu	Ile	Gly	Lys 125	Ser	Pro	Phe
Суѕ	Glu 130	Leu	Asn	Arg	Ser	Glu 135	Ile	Ala	Gly	Ile	11e 140	Arg	Glu	Val	Thr
Thr 145		Arg	Arg	Phe	Cys 150	Met	Tyr	Tyr		Lys 155		Val	Trp	Asn	Ile 160
His	Leu	Glu	Thr	Gly 165	Ile	Pro	Pro	Ala	Asn 170	Trp	Ala	Lys	Lys	Gly 175	Phe
Asn	Glu	Asn	Glu 180	Lys	Phe	Ala	Ala	Phe 185		Phe	Phe	Leu	Gly 190	.Val	Thr
Asp	Glu	Ser 195		Leu	Glu	Pro	Lys 200		Gly	Val	Lys	Arg 205		Pro	Thr
Lys	Ala 210				· .		•								

The DNA molecule which constitutes a substantial portion of the RSPaV strain RSP158 genome comprises the nucleotide sequence corresponding to SEQ. ID. No. 23 as follows:

					•	
GAAGCTAGCA	CATTTCTGTT	CAACACTATG	GCTAACATGT	TGTTCACTTT	TCTGAGATAT	60
GAACTGACGG	GTTCAGAGTC	AATAGCATTT	GCAGGGGATG	ATATGTGTGC	TAATAGAAGG	120
TTGCGGCTTA	AAACGGAGCA	TGAGGGTTTT	CTGAACATGA	TCTGCCTTAA	GGCCAAGGTT	180
CAGTTTGTTT	CCAACCCCAC	ATTCTGTGGA	TGGTGCTTAT	TTAAGGAGGG	AATCTTCAAG	240
AAACCTCAAC	TAATTTGGGA	GCGAATATGC	ATAGCCAGAG	AGATGGGCAA	TCTGGAGAAC	300
TGTATTGACA	ATTATGCGAT	AGAAGTGTCC	TATGCATATA	GATTGGGTGA	GCTATCAATT	360
GAAATGATGA	CAGAAGAAGA	AGTGGAGGCA	CACTACAATT	GTGTGAGGTT	CCTGGTTAGG	420
AACAAGCATA	AGATGAGGTG	CTCAATTTCA	GGCCTGTTTG	AAGTGGTTGA	TTAGGCCTTA	480
AGTATTTGGC	GTTGTTCGAG	TTATTATGAA	TAATTTAGTT	AAAGCATTAT	CAGCCTTCGA	540
GTTTATAGGT	GTTTTCAATG	TGCTCAAATT	TCCAGTTGTT	ATACATAGTO	TGCCTGGTAG	600
TGGTAAGAGT	AGCTTAATAA	GGGAATTAAT	CTCAGAGGAC	GAGAGTTTC	TGGCTTTCAC	660
AGCAGGTGTT	CCAGACAGTO	CTAACCTCAC	AGGGAGGTAG	C ATCAAGCCT	r ACTCACCAGG	720
ATGCGCAGT	CAAGGAAAA	G TGAATATAC	TGATGAGTA	C TTGTCCGTT	C AAGACATTTC	780
GGGTTTTGAT	GTACTGTTT	r CAGACCCGT	A CCAGAATAT	C AGTATTCCC	C AAGAGGCGCA	840
TTTCATTAA	TCCAAGACT	r gtaggtttg	G TGTGAACAC	T TGCAAATAC	C TTTCCTCTTT	900
CGGTTTCGA	A GTTAGCAGC	G ACGGGCTGG	A CGACGTCAT	T GTGGGATCG	C CCTTCACTCT	960

AGATGTTGAA	GGGGTGCTGA	TATGTTTTGG	CAAGGAGGCG	GTAGATCTCG	CTGTTGCGCA	1020
CAACTCTGAA	TTCAAGTTGC	CGTGTGAGGT	TCGAGGTTCA	ACCTTCAATG	TGGTAACCCT	1080
TTTGÄAATCA	AGAGACCCAA	CCCCAGAGGA	CAGGCACTGG	TTTTACATCG	CTGCCACAAG	1140
ACATAGGAAG	AAATTGGTCA	TTATGCAGTA	AAATGCCTTT	TCAGCAGCCT	GCTAATTGGG	1200
CAAAAACCAT	AACTCCATTG	ACTATTGGCT	TAGGAATTGG	ACTTGTGCTG	CATTTTCTGA	1260
GAAAGTCAAA	TCTACCATAT	TCAGGAGACA	ACATCCATCA	ATTTCCTCAC	GGGGGCGTT	1320
ACCGGGACGG	CACAAAAAGT	ATAACTTACT	GTGGCCCTAA	GCAGTCCTTC	CCCAGTTCAG	1380
GAATATTTGG	TCAGTCTGAG	AATTTTGTGC	CCTTAATGCT	TGTCATAGGT	CTAATTGCAT	1440
TCATACATGT	ATTGTCTGTT	TGGAATTCTG	GTCTTGGTAG	GAATTGCAAT	TGCCATCCAA	1500
ATCCTTGCTC	ATGTAGACAA	CAGTAGTGGC	AGTCACCAAG	GTTGCTTTAT	CAGGGCCACT	1560
GGAGAGTCTA	TTTTGATTGA	AAATTGTGGC	CCAAGCGAGG	CCCTTGCATC	AACAGTGAGG	1620
GAGGTGTTGG	GGGGTTTGAA	GGCTTTAGGA	ATTAGCCATA	CTACTGAAGA	AATTGATTAT	1680
CGTTGTTAAA	TTGGTTAAAT	GGCGAGTCAA	GTTGGTAAGC	TCCCCGGAGA	ATCAAATGAG	1740
GCATTTGAAG	CCCGGCTGAA	ATCACTGGAG	TTGGCTAGAG	CTCAAAAGCA	GCCAGAAGGT	1800
TCAAACACAC	CGCCTACTCT	CAGTGGTGTG	CTTGCCAAAC	GTAAGAGGGT	TATTGAGAAT	1860
GCACTCTCAA	AGACAGTGGA	CATGAGGGAG	GTGTTGAAAC	ACGAAACGGT	TGTAATTTCC	1920
CCAAATGTCA	TGGATGAGGG	TGCAATAGAT	GAACTGATTO	GTGCATTCG	S AGAATCAGGC	1980
ATAGCTGAGA	GCGCACAATT	TGATGTGGC		.*		2009

The RSP158 strain contains five open reading frames (i.e., ORF1-5). ORF1 and ORF5 are only partially sequenced. The nucleotide sequence of RSP158 is 87.6% identical to the corresponding region of RSPaV-1 (type strain). The numbers of amino acid residues of corresponding ORFs of RSP158 and RSPaV-1 (type strain) are exactly the same. In addition, the amino acid sequences of these ORFs have high identities to those of RSPaV-1: 99.3% for ORF1, 95% for ORF2, 99.1% for ORF3, 88.8% for ORF4, and 95.1% for ORF5. The nucleotide and amino acid sequence information of the RSP158 ORFs are described below.

Another DNA molecule of the present invention (RSP158 incomplete ORF1)

includes nucleotides 1-447 of SEQ. ID. No. 23. This DNA molecule is believed to code for a polypeptide portion of a RSP158 replicase and comprises a nucleotide sequence corresponding to SEQ. ID. No. 24 as follows:

GAAGCTAGCA	CATTTCTGTT	CAACACTATG	GCTAACATGT	TGTTCACTTT	TCTGAGATAT	60
GAACTGACGG	GTTCAGAGTC	AATAGCATTT	GCAGGGGATG	ATATGTGTGC	TAATAGAAGG	120
TTGCGGCTTA	AAACGGAGCA	TGAGGGTTTT	CTGAACATGA	TCTGCCTTAA	GGCCAAGGTT	180
CAGTTTGTTT	CCAACCCCAC	ATTCTGTGGA	TGGTGCTTAT	TTAAGGAGGG	AATCTTCAAG	240
AAACCTCAAC	TAATTTGGGA	GCGAATATGC	ATAGCCAGAG	AGATGGGCAA	TCTGGAGAAC	300
TGTATTGACA	ATTATGCGAT	AGAAGTGTCC	TATGCATATA	GATTGGGTGA	GCTATCAATT	360
GAAATGATGA	CAGAAGAAGA	AGTGGAGGCA	CACTACAATT	GTGTGAGGTT	CCTGGTTAGG	420
AACAAGCATA	AGATGAGGTG	CTCAATT				44

The polypeptide encoded by the nucleotide sequence of SEQ. ID. No. 24 has a deduced amino acid sequence corresponding to SEQ. ID. No. 25 as follows:

Glu 1	Ala	Ser	Thr	Phe 5	Leu	Phe	Asn	Thr	Met 10	Ala	Asn	Met	Leu	Phe 15	Thr
Phe	Leu	Arg	Tyr 20	Glu	Leu	Thr	Gly	Ser 25	Glu	Ser	Ile	Ala	Phe 30	Ala	Gly
Asp	Asp	Met 35	Cys	Ala	Asn	Arg	Arg 40	Leu	Arg	Leu	Lys	Thr	Glu	His	Glu
Gly	Phe 50	Leu	Asn	Met	Ile	Cys 55	Leu	Lys	Ala	Lys	Val 60	Gln	Phe	Val	Ser
Asn 65	Pro	Thr	Phe	Cys	Gly 70	Trp	Cys	Leu	Phe	Lys 75	Glu	Gly	Ile	Phe	Lys 80
Lys	Pro	Gln	Leu	Ile 85	Trp	Glu	Arg	Ile	Cys 90	Ile	Ala	Arg	Glu	Met 95	Gly
Asn	Leu	Glu	Asn 100		: Ile	Asp	Asn	105	Ala	Ile	e Glu	Val	Ser 110	Tyr	Ala
Tyr	Arg	Leu 115		Glu	l Leu	Ser	11e		ı Met	: Met	Thi	Glu 125	Glu S	Glu	Val
Glu	130		5 Туг	Ası	ı Cys	Val		g Phe	e Le	ı Val	1 Arg	g Asr	ı Lys	s His	Lys
Met	Arç	g Cys	s Sei	r Ile	е										

Another DNA molecule of the present invention (RSP158 ORF2) includes nucleotides 506-1171 of SEQ. ID. No. 23. This DNA molecule codes for a first protein or polypeptide of the RSP158 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 26 as follows:

ATGAATAATT	TAGTTAAAGC	ATTATCAGCC	TTCGAGTTTA	TAGGTGTTTT	CAATGTGCTC	60
AAATTTCCAG	TTGTTATACA	TAGTGTGCCT	GGTAGTGGTA	AGAGTAGCTT	AATAAGGGAA	120
TTAATCTCAG	AGGACGAGAG	TTTCGTGGCT	TTCACAGCAG	GTGTTCCAGA	CAGTCCTAAC	180
CTCACAGGGA	GGTACATCAA	GCCTTACTCA	CCAGGATGCG	CAGTGCAAGG	AAAAGTGAAT	240
ATACTTGATG	AGTACTTGTC	CGTTCAAGAC	ATTTCGGGTT	TTGATGTACT	GTTTTCAGAC	300
CCGTACCAGA	ATATCAGTAT	TCCCCAAGAG	GCGCATTTCA	TTAAGTCCAA	GACTTGTAGG	360
TTTGGTGTGA	ACACTTGCAA	ATACCTTTCC	TCTTTCGGTT	TCGAAGTTAG	CAGCGACGGG	420
CTGGACGACG	TCATTGTGGG	ATCGCCCTTC	ACTCTAGATG	TTGAAGGGGT	GCTGATATGT	480
TTTGGCAAGG	AGGCGGTAGA	TCTCGCTGTT	GCGCACAACT	CTGAATTCAA	GTTGCCGTGT	540
GAGGTTCGAG	GTTCAACCTT	CAATGTGGTA	ACCCTTTTGA	AATCAAGAGA	CCCAACCCCA	600
GAGGACAGGC	ACTGGTTTTA	CATCGCTGCC	ACAAGACATA	GGAAGAAATT	GGTCATTATG	660
CAGTAA				•		666

The first protein or polypeptide of the RSP158 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 27 as follows:

Met 1	Asn	Asn	Leu	Val 5	гàг	Ala	Leu	Ser	Ala 10	Phe	GLu	Phe	Ile	Gly 15	Val
Phe	Asn	Val	Leu 20	Lys	Phe	Pro	Val	Val 25	Ile	His	Ser	Val	Pro 30	Gly	Ser
Gly	Lys	Ser 35	Ser	Leu	Ile	Arg	Glu 40	Leu	Ile	Ser	Glu	Asp 45	Glu	Ser	Phe
Val	Ala 50	Phe	Thr	Ala	Gly	Val 55	Pro	Asp	Ser	Pro	Asn 60	Leu	Thr	Gly	Arg
Tyr 65	Ile	Lys	Pro	Tyr	Ser 70	Pro	Gly	Cys	Ala	Val 75	Gln	Gly	Lys	Val	Asn 80
Ile	Leu	Asp	Glu	Tyr 85	Leu	Ser	Val	Gln	Asp 90	Ile	Ser	Gly	Phe	Asp 95	Val
Leu	Phe	Ser	Asp 100	Pro	Tyr	Gln	Asn	Ile 105	Ser	Ile	Pro	Gln	Glu 110	Ala	His
Phe	Ile	Lys 115	Ser	Lys	Thr	Cys	Arg 120		Gly	Val	Asn	Thr 125	_	Lys	Tyr
Leu	Ser 130	Ser	Phe	Gly	Phe	Glu 135	Val	Ser	Ser	Asp	Gly 140		Asp	Asp	Val
11e 145		Gly	Ser	Pro	Phe 150		Leu	Asp	Val	Glu 155	_	Val	Leu	Ile	Cys 160

Phe	Gly	Lys	Glu	Ala 165	Val	Asp	Leu	Ala	Val 170	Ala	His	Asn	Ser	Glu 175	Phe
Lys ·	Leu	Pro	Cys 180	Glu	Val	Arg	Gly	Ser 185	Thr	Phe	Asn	Val	Val 190	Thr	Leu
Leu	Lys	Ser 195	Arg	Asp	Pro	Thr	Pro 200	Glu	Asp	Arg	His	Trp 205	Phe	Tyr	Ile
Ala	Ala 210	Thr	Arg	His	Arg	Lys 215	Lys	Leu	Val	Ile	Met 220	Gln			

The first protein or polypeptide of the RSP158 triple gene block has a molecular weight of about 20 to 26 kDa., preferably 24.4 kDa.

Another DNA molecule of the present invention (RSP158 ORF3) includes nucleotides 1173-1526 of SEQ. ID. No. 23. This DNA molecule codes for a second protein or polypeptide of the RSP158 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 28 as follows:

ATGCCTTTTC	AGCAGCCTGC	TAATTGGGCA	AAAACCATAA	CTCCATTGAC	TATTGGCTTA		60
GGAATTGGAC	TTGTGCTGCA	TTTTCTGAGA	AAGTCAAATC	TACCATATTC	AGGAGACAAC		120
ATCCATCAAT	TTCCTCACGG	GGGGCGTTAC	CGGGACGGCA	CAAAAAGTAT	AACTTACTGT		180
GGCCCTAAGC	AGTCCTTCCC	CAGTTCAGGA	ATATTTGGTC	AGTCTGAGAA	TTTTGTGCCC		240
TTAATGCTTG	TCATAGGTCT	AATTGCATTC	ATACATGTAT	TGTCTGTTTG	GAATTCTGGT	• • •	300
CTTGGTAGGA	ATTGCAATTG	CCATCCAAAT	CCTTGCTCAT	GTAGACAACA	GTAG		354

The second protein or polypeptide of the RSP158 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 29 as follows:

Met 1	Pro	Phe	Gln	Gln 5	Pro	Ala	Asn	Trp	Ala 10	Lys	Thr	Ile	Thr	Pro 15	Leu
Thr	Ile '	Gly	Leu 20	Gly	Ile	Gly	Leu	Val 25	Leu	His	Phe	Leu	Arg 30	Lys	Ser
Asn	Leu	Pro 35	Tyr	Ser	Gly	Asp	Asn 40	Ile	His	Gln	Phe	Pro 45	His	Gly	Gly
Arg	Tyr 50	Arg	Asp	Gly	Thr	Lys 55	Ile	Thr	Tyr	Суѕ	Gly 60	Pro	Lys	Gln	Ser
Phe 65	Pro	Ser	Ser	Gly	Ile 70	Phe	Gly	Gln	Ser	Glu 75	Asn	Phe	Val	Pro	Leu 80
Met	Leu	Val	Ile	Gly 85	Leu	Ile	Ala	Phe	Ile 90	His	Val	Leu	Ser	Val 95	Trp

Asn Ser Gly Leu Gly Arg Asn Cys Asn Cys His Pro Asn Pro Cys Ser
100 105 110

Cys Arg Gln Gln
115

The second protein or polypeptide of the RSP158 triple gene block has a molecular weight of about 10 to 15 kDa., preferably 12.9 kDa.

Another DNA molecule of the present invention (RSP158 ORF4) includes nucleotides 1447-1689 of SEQ. ID. No. 23. This DNA molecule codes for a third protein or polypeptide of the RSP158 triple gene block and comprises a nucleotide sequence corresponding to SEQ. ID. No. 30 as follows:

5

10

TAA						243
TTGGGGGGTT	TGAAGGCTTT	AGGAATTAGC	CATACTACTG	AAGAAATTGA	TTATCGTTGT	240
TCTATTTTGA	TTGAAAATTG	TGGCCCAAGC	GAGGCCCTTG	CATCAACAGT	GAGGGAGGTG	180
GCTCATGTAG	ACAACAGTAG	TGGCAGTCAC	CAAGGTTGCT	TTATCAGGGC	CACTGGAGAG	120
ATGTATTGTC	TGTTTGGAAT	TCTGGTCTTG	GTAGGAATTG	CAATTGCCAT	CCAAATCCTT	60

The third protein or polypeptide of the RSP158 triple gene block has a deduced amino acid sequence corresponding to SEQ. ID. No. 31 as follows:

Met 1	Tyr	Cys	Leu	Phe 5	Gly	Ile	Leu	Val	Leu 10		Gly	Ile	Ala	Ile 15	Ala
Ile	Gln	Ile	Leu 20	Ala	His	Val	Asp	Asn 25	Ser	Ser	Gly.		His 30	Gln	Gly
Cys	Phe	11e 35	_	Ala	Thr	Gly	Glu 40	Ser	Ile	Leu	Ile	Glu 45	Asn	Cys	Gly
Pro	Ser 50	Glu	Ala	Leu	Ala	Ser 55	Thr	Val	Arg	Glu	Val 60	Leu	Gly	Gly	Leu
Lys	Ala	Leu	Gly	Ile	Ser	His	Thr	Thr	Glu	Glu 75	Ile	Aşp	Tyr	Arg	Cys 80

The third protein or polypeptide of the RSP158 triple gene block has a molecular weight of about 5 to 10 kDa., preferably 8.4 kDa.

Yet another DNA molecule of the present invention (RSP158 ORF5) includes nucleotides 1699-2009 of SEQ. ID. No. 23. This DNA molecule codes for a partial RSP158 coat protein or polypeptide and comprises a nucleotide sequence corresponding to SEQ. ID. No. 32 as follows:

ATGGCGAGTC	AAGTTGGTAA	GCTCCCCGGA	GAATCAAATG	AGGCATTTGA	AGCCCGGCTG	60
AAATCACTGG	AGTTGGCTAG	AGCTCAAAAG	CAGCCAGAAG	GTTCAAACAC	ACCGCCTACT	120
CTCAGTGGTG	TGCTTGCCAA	ACGTAAGAGG	GTTATTGAGA	ATGCACTCTC	AAAGACAGTG	180
GACATGAGGG	AGGTGTTGAA	ACACGAAACG	GTTGTAATTT	CCCCAAATGT	CATGGATGAG	240
GGTGCAATAG	ATGAACTGAT	TCGTGCATTC	GGAGAATCAG	GCATAGCTGA	GAGCGCACAA	300
TTTGATGTGG	С					313

The polypeptide has a deduced amino acid sequence corresponding to SEQ. ID. No. 33 as follows:

Met Ala Ser Gln Val Gly Lys Leu Pro Gly Glu Ser Asn Glu Ala Phe 15

Glu Ala Arg Leu Lys Ser Leu Glu Leu Ala Arg Ala Gln Lys Gln Pro 25

Glu Gly Ser Asn Thr Pro Pro Thr Leu Ser Gly Val Leu Ala Lys Arg 45

Lys Arg Val Ile Glu Asn Ala Leu Ser Lys Thr Val Asp Met Arg Glu 50

Val Leu Lys His Glu Thr Val Val Ile Ser Pro Asn Val Met Asp Glu 65

Gly Ala Ile Asp Glu Leu Ile Arg Ala Phe Gly Glu Ser Gly Ile Ala 95

Glu Ser Ala Gln Phe Asp Val

The following seven cDNA clones are located at the central part of the ORF1 of RSPaV-1 and all have high identities (83.6-98.4%) in nucleotide sequence with the comparable regions of RSPaV-1. When their nucleotide sequences are aligned with MegAlign (DNAStar), a highly conserved region of ca. 600 nucleotides was found. The universal primers BM98-3F/BM98-3R (SEQ. ID. Nos. 51 and 52, infra) were designed based on the conserved nucleotide sequences of this region.

Portions of the genome from yet other strains of *Rupestris* stem pitting associated viruses have also been isolated and sequenced. These include strains designated 140/94-19 (T7+R1), 140/94-24 (T7+R1), 140/94-2 (T3+F1), 140/94+42 (T7+R1), 140/94-64 (T7+R1), 140-94-72 (T7+R1), and 140/94-6 (T3+BM98-3F+F2).

The nucleotide sequence of 140/94-19 (T7+R1) corresponds to SEQ. ID. No. 34 as follows:

GCAGGATTGA	AGGCTGGCCA	CTGTGTGATT	TTTGATGAGG	TCCAGTTGTT	TCCTCCTGGA	60
TACATCGATC	TATGCTTGCT	TATTATACGT	AGTGATGCTT	TCATTTCACT	TGCCGGTGAT	120
CCATGTCAAA	GCACATATGA	TTCGCAAAAG	GATCGGGCAA	TTTTGGGCGC	TGAGCAGAGT	180
GACATACTTA	GAATGCTTGA	GGGCAAAACG	TATAGGTATA	ACATAGAAAG	CAGGAGGTTT	240
GTGAACCCAA	TGTTCGAATC	AAGACTGCCA	TGTCACTTCA	AAAAGGGTTC	GATGACTGCC	300
GCTTTCGCTG	ATTATGCAAT	CTTCCATAAT	ATGCATGACT	TTCTCCTGGC	GAGGTCAAAA	360
GGTCCTTTGG	ATGCCGTTTT	GGTTTCCAGT	TTTGAGGAGA	AAAAGATAGT	CCAGTCCTAC	420
TTTGGAATGA	AACAGCTCAC	ACTCACATTT	GGTGAATCAA	CTGGGTTGAA	TTTCAAAAAT	480
GGGGGAATTC	TCATATCACA	TGATTCCTTT	CACACAGATG	ATCGGCCGGT	GGCTTACTGC	540
TTTATCTCGC	TTCAGCCACA	ATTTGGATTT	GGTGAACATT	ACAGGTCTGA	GGGTGGAAAG	600
TTTCCTCTCG	CACTTTGCTG	GCAAACCCCT	CTACCATTTT	TTAACAGCCA	AAAGTGGGGA	660
GAATGTCATA	CGAGATTTGC	TCCCAGGTGA	GCCTAACTTC	TTCAGTGGCT	TTAACGTTAG	720
CATTGGAAAG	AATGAAGGTG	TTAGGGAGGA	GAAGTTATGT	GGTGACCCAT	GGTTAAAAGT	780
CATGCTTTTC	CTGGGTCAAG	ATGAGGATTG	TGAAGTTGAA	GAGATGGAGT	CAGAGTGCTC	840
AAATGAAGAA	TGGTTTAAAA	CCCACATTCC	CCTGAGTAAT	CTGGAGTCAA	CCAGGGCTAG	900
GTGGGTGGGT	AAAATGGCTT	TGAAAGAGTA	TCGGGAGGTG	GCTTGTGGTT	ATGAAATGAC	960
TCAACAATTO	: TTTGATGAGC	ATAGGGGTGG	AACTGGTGA	CAACTGAGC	ATGCATGTGA	1020
GAGGTTTGA	AGCATTTACC	CAAGGCATA	A AGGAAATGA	TCAATAACC	TCCTTATGGC	1080
TGTCCGAAA	G CGTCTCAAAT	TTTCGAAGC	CCAGGTTGA	A GCTGCCAAA	TGAGGCGGC	1140
CAAACCATAT	GGGAAATTCT	TATTAGACT	TCCTATCCA	A AATCCCATTO	AAAGCCAGTC	1200
TTAATA						1206

The nucleotide sequence of 140/94-24 (T7+R1) corresponds to SEQ. ID. No. 35 as follows:

60	CATAAGGCAT	CCAATGAGCA	TTTGATATAA	TTCCGCCTTG	ATGGTAAGAT	ATTAACCCAA
120	GAAGGTTAAG	ATGAGTTGAG	ATAAAAGTAG	CCCTCAGAGC	TCGGCAATGG	GTTGAGAAGA
180	TCCAAACTTT	TAACCTATTT	GGGTCCAAAA	TTCAATGAAT	TTGATCTTCT	CGATCCGCCC
240	CATAAGTGAT	TAACTGGTGT	CTAGGGGGCC	AGGGTGCTTG	AAAAGTTGCA	GAGCGGGCTG

GAAAAGTTCA	GTGATGCAAA	ACCCTGGCTT	TCTGGTATAT	CAACTGCGGA	TATAAAGCCA	300
AGAGAGCTAA	CTGTCGTGCT	TGGCACTTTT	GGGGCTGGAA	AGAGTTTCTT	GTATAAGAGT	360
TTCATGAAGA	GATCTGAGGG	AAAATTTGTA	ACTTTTGTTT	CCCCTAGACG	AGCCTTGGCA	420
AATTCAATCA	AAAATGATCT	TGAAATGGAT	GATGGCTGCA	AAGTTGCCAA	AGCAGGCAAA	480
TCAAAGAAGG	AAGGGTGGGA	TGTAGTGACC	TTTGAAGTTT	TCCTTAGAAA	AGTTTCTGGT	540
TTGAAAGCTG	GTCATTGTGT	GATTTTTGAT	GAGGTTCAGT	TGTTTCCCCC	TGGATACATC	600
GATCTGTGTT	TACTTGTCAT	ACGAAGTGAT	GCTTTCATTT	CACTTGCTGG	TGATCCATGC	660
CAGAGCACAT	ATGATTCACA	GAAGGATCGA	GCAATTTTGG	GAGCTGAGCA	GAGTGACATA	720
CTCAGACTGC	TTGAAGGAAA	GACATATAGG	TACAACATAG	AAAGCAGACG	TTTTGTGAAC	780
CCAATGTTTG	AATCTAGACT	ACCATGTCAC	TTCAAAAAGG	GTTCAATGAC	TGCAGCCTTT	840
GCTGATTATG	CAATCTTCCA	CAATATGCAT	GACTTCCTCC	TGGCGAGGTC	AAAAGGCCCC	900
TTGGATGCTG	TTCTAGTTTC	CAGTTTTGAG	GAĞAAGAAA	TAGTCCAATC	CTACTTTGGG	- 960
ATGAAGCAAC	TCACTCTCAC	: ATTTGGTGAF	TCAACTGGGT	TGAACTTCA	AAATGGAGGA	1020
ATTCTCATAT	CACATGACTO	CTTTCATACT	GACGATCGA	GGTGGCTTAG	C TGCTTTATCT	1080
CGATTCAGC	: ATAATTTGG	A TTTGGTGAA	C ATCACAGGT	TTGAGGGTG	G AAAGTTTTCT	1140
CTCACATTT	GCTGGTAAA	CCCTTTACC	A CTTTTTGAC	G GCTTAAAAG	T GGAGAGAATG	1200
TCATACGAGA	A CCTGCTTCA	GTGAGCCTA	A CTTCTTTA	G GGGTTCAAT	G TCAGCATTGG	- 1260
AAAAAAATG	AAGGGGTTA	G AGAA			,	1284

The nucleotide sequence of 140/94-2 (T3+F1) corresponds to SEQ. ID. No. 36 as follows:

CATTTTTAAA	ATTTAATCCA	GTCGACTCAC	CAAATGTGAG	CGTAAGCTGT	TTCATCCCAA	60
AGTAGGACTG	GACTATTTTC	TTCTCCTCAA	AACTAGAAAC	CAGAATGGCA	TCCAAAGGAC	120
CTTTTGACCT	TGCCAGGAGG	AAATCATGCA	TATTGTGGAA	AATGGCATAA	TCAGCAAAGG	180
CAGCAGTCAT	TGTACCCTTT	TTGAAGTGAC	ATGGCAGTCG	AGATTCAAAC	ATTGGGTTCA	240
CAAATCTTCT	GCTTTCTATG	TTGTACCTAT	ACGTCTTGCC	TTCAAGTATT	TTGAGTATGT	300
CACTCTGCTC	AGCGCCCAAA	ATCGCCCGAT	CTTTTTGTGA	GTCATATGTG	CTCTGACATG	360
GGTCACCAGC	AAGTGAAATG	AAAGCATCAC	TACGTATAAT	AAGCAAACAT	AGATCGATGT	420
ATCCAGGGGG	AAACAACTGG	ACCTCATCGA	AAATTACACA	GTGACCAGCT	TTTAGACCTG	480
CAACTTTTCT	AAGGAAGACT	TCAAAAGTCA	CAACATCCCA	TCCTTCCTTC	TTTGACCTGC	540
CTGCTTTGGC	AACTTTGCAG	CTATCATCCA	TTTCAAGATC	ATTTTTGATT	GAATTCGCTA	600

GAGCCCGTCT	GGGGGAAACA	AAAGTTACGA	ATTTACCCTC	AGATCTTTTC	ATAAAGCTCT	660
IGTACAAAAA	GCTTTTTCCG	GCTCCAAATG	TGCCAAGCAC	AACAGTTAGC	TCCCTCGGCT	720
TAATGTCAGT	AGTTGATATA	CCAGAAAGCC	AGGGCTTTGC	ATCACTGAAC	TTCTCATCAC	780
TTATGACACC	AGTTAGGCCT	CCTAGCAGAC	ACCCTTGCAA	CTTTTCAGCC	CGCTCAAAAC	840
TTGGGAAGTA	GGTTACCTTG	GACCCATTAA	TTGAAAGAAG	ATCAAGGGCG	GATCGCTTGA	900
CCTTTCGCAA	TTCATCTACT	TTAATGCTCT	GAGGGCCATT	ACCTATCTTT	TCAACATGCC	960
TTATGTGCTC	ATTAGTTATG	TCAAACAGAG	CGGAAAACTT	GCCATGTGGA	TTAATCACCT	1020
CAATTTCCCC	ATTTATGTCA	CACTTAGCGC	AAATGTCAAA	AGCCTCAAAG	GCTTCAGCTA	1080
AGTTACATCA	TGTTGAGCCT	CCCCCTTGGC	AAAGCTCCTC	AAAAATGTGG	TTAGTGCTAG	1140
GCCTGCACAA	TAATTAACAC	ATCAACTTCA	CCCTGCCAAT	GCTGAACAAT	ACTGTTATCA	1200
TGCAACCATC	CATGGGGCAC	ATGGTTGGAA	TTGATTGATT	TAAGGCAAAA	ATCCCCACAG	1260
GGGGCATCCC	CTTCCCCAAT	TTCCACTGAT	TCATACTCTG	GCGTTATCAT	ATCAACCCAA	1320
TGTGTCAAAT	ACAAATAATG	CAATCTCTCA	TCTCCGATAA	CATTTCCCCC	AAATTTTTAAA	1380
AATGGTGGGG	TGAAAATTGG	AA				1402

The nucleotide sequence of 140/94-42 (T7+R1) corresponds to SEQ. ID. No. 37 as follows:

GTGGTTTTTG	CAACAACAGG	CCCAGGTCTA	TCTAAGGTTT	TGGAAATGCC	TCGAAGCAAG	60
AAGCAATCTA	TTCTGGTTCT	TGAGGGAGCC	CTATCCATAG	AAACGGACTA	TGGCCCAAAA	120
GTTCTGGGAT	CTTTTGAAGT	TTTCAAAGGG	GATTTCAACA	TTAAAAAAAT	GGAAGAAAGT	180
TCCATCTTTG	TAATAACATA	CAAGGCCCCA	GTTAGATCTA	CTGGCAAGTT	GAGGGTCCAC	240
CAATCAGAAT	GCTCATTTTC	TGGATCCAAG	GAGGTATTGC	TGGGTTGTCA	GATTGAGGCA	300
TGTGCTGATT	ATGATATTGA	TGATTTCAAT	ACTTTCTTTG	TACCTGGTGA	TGGTAATTGC	360
TTTTGGCATT	CAGTTGGTTT	CTTACTCAGT	ACTGACGGAC	TTGCTTTGAA	GGCCGGCATT	420
CGTTCTTTCG	TGGAGAGTGA	ACGCCTGGTG	AGTCCAGATC	TTTCAGCCCC	AACCATTTCT	480
AAACAACTGG	GGGAAAATGC	TTATGCCGAG	AATGAGATGA	TTGCATTATT	TTGTATTCGA	540
CACCATGTGA	GGCTGATAGT	GATTACGCCA	GAGTATGAAG	TCAGTTGGAA	ATTTGGGGAA	600
GGTGAATGGC	CCCTGTGCGG	AATTCTTTGC	СТТАААТСАА	ATCACTTCCA	ACCATGTGCC	660
CCATTGAATG	GTTGCATGAT	TACAGCTATT	GCTTCAGCAC	TTGGTAGGCG	TGAAGTTGAT	720
GTGCTTAATT	ATCTGTGCAG	GCCTAGCACT	AACCACATTT	TTGAGGAGCT	TTGCCAAGGG	780

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GGAGGCCTCA	ACATGATGTA	CTTAGCTGAA	GCCTTTGAGG	CTTTTGACAT	TTGCGCTAAG	840
TGTGACATAA	ATGGGGAAAT	TGAGGTGATT	AATCCACATG	GCAAGTTTTC	CGCTCTGTTT	900
GACATAACTA	ATGAGCACAT	AAGGCATGTT	GAAAAGATAG	GTAATGGCCC	TCAGAGCATT	960
AAAGTAGATG	AATTGCGAAA	GGTCAAGCGA	TCTGCCCTTG	ATCTTCTTTC	AATTAATGGG	1020
TCCAAGGTAA	CCTACTTCCC	AAGTTTTGAG	CGGGCTGAAA	AGTTGCAAGG	GTGTCTGCTA	1080
GGAGGCCTAA	CTGGTGTCAT	AAGTGATGAG	AAAGTCAGTG	ATGCAAAGCC	CTGCTTTTTG	1140
GTATATCAAC	TACTGACATT	AAGCCGAGGG	AGCTAACTGT	TGTGCTTTGG	CACATTTGGA	1200
GCCCGGAAAA	AGCCTTTTGT	ACCAAGAGCT	TTATTG	· .		123

The nucleotide sequence of 140/94-6 (T3 + BM98 - 3F + F2) corresponds to SEQ. ID. No. 38 as follows:

GTCTAACTGG CGTTATAAGT GATGAGAAAT TCAGTGATGC AAAACCTTGG CTTTCTGGTA TATCTACTAC AGATATTAAG CCAAGGGAAT TAACTGTTGT GCTTGGTACA TTTGGGGCTG GGAAGAGTTT CTTGTACAAG AGTTTCATGA AAAGGTCTGA GGGTAAATTC GTAACCTTTG TTTCTCCCAG ACGTGCTTTA GCAAATTCAA TCAAAAATGA TCTTGAAATG GATGATAGCT GCAAAGTTGC CAAAGCAGGT AGGTCAAAGA AGGAAGGGTG GGATGTAGTA ACTTTTGAGG TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTT GATGAGGTCC AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT TGGGCGCTGA GCAGAGTGAC ATACTTAGAA TGCTTGAGGG CAAAACGTAT AGGTATAACA	
GGAAGAGTTT CTTGTACAAG AGTTTCATGA AAAGGTCTGA GGGTAAATTC GTAACCTTTG TTTCTCCCAG ACGTGCTTTA GCAAATTCAA TCAAAAATGA TCTTGAAATG GATGATAGCT GCAAAGTTGC CAAAGCAGGT AGGTCAAAGA AGGAAGGGTG GGATGTAGTA ACTTTTGAGG TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTTT GATGAGGTCC AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	60
TTTCTCCCAG ACGTGCTTTA GCAAATTCAA TCAAAAATGA TCTTGAAATG GATGATAGCT GCAAAGTTGC CAAAGCAGGT AGGTCAAAGA AGGAAGGGTG GGATGTAGTA ACTTTTGAGG TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTTT GATGAGGTCC AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	120
GCAAAGTTGC CAAAGCAGGT AGGTCAAAGA AGGAAGGGTG GGATGTAGTA ACTTTTGAGG TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTTT GATGAGGTCC AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	1 80
TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTT GATGAGGTCC AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	240
AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	300
TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT	360
	420
TGGGCGCTGA GCAGAGTGAC ATACTTAGAA TGCTTGAGGG CAAAACGTAT AGGTATAACA	480
	540
TAGAAAGCAG GAGGTTTGTG AACCCAATGT TCGAATCAAG ACTGCCATGT CACTTCAAAA	600
AGGGTTCGAT GACTGCCGCT TTCGCTGATT ATGCAATCTT CCATAATATG CATGACTTTC	660
TCCTGGCGAG GTCAAAAGGT CCTTTGGATG CCGTTTTGGT TTCCAGTTTT GAGGAGAAAA	720
AGATAGTCCA GTCCTACTTT GGAATGAAAC AGCTCACACT CACATTTGGT GAATCAACTG	780
GGTTGAATTT CAAAAATGGG GGAATTCTCA TATCACATGA TTCCTTTCAC ACAGATGATC	840
GGCGGTGGCT TACTGCTTTA TCTCGCTTCA GCCACAATTT GGATTTGGTG AACATTACAG	900
GTCTGAGGTG GAAAGTTTCC TCTCGCACTT TGCTGGCAAA CCCCTCTACC ATTTTTTAAC	960
AGCCAAAAGT GGGGAGAATG TCATACGAGA TTTGCTCCCA GGTGAGCCTA ACTTCTTCAG	1020
TGGCTTTAAC GTTAGCATTG GAAAGAATGA AGGTGTTAGG GAGGAGAAGT TATGTGGTGA	1080
CCCATGGTTA AAAGTCATGC TTTTCCTGGG TCAAGATGAG GATTGTGAAG TTGAAGAGAT	1140
GGAGTCAGAG TGCTCAAATG AAGAATGGTT TAAAACCCAC ATTCCCCTGA GTAATCTGGA	1200

GTCAACCAGG	GCTAGGTGGG	TGGGTAAAAT	GGCCTTGAAA	GAGTATCGGG A	AGGTGCGTTG	1260
TGGTTATGAA	ATGACTCAAC	AATTCTTTGA	TGACAT			1296
	The nucleot	ide sequence	of 140/94-64 (T7+R1) corres	ponds to SEQ.	
ID. No. 39 as	follows:	-	·	·	-	
ATGTTCACCA	AATCCAAATT	ATGGCTGAAG	CGAGATAAAG	CAGTAAGCCA (CCGCCGATCA	60
TCTGTGTGAA	AGGAATCATG	TGATATGAGA	ATTCCCCCAT	TTTTGAAATT (CAACCCAGTT	120
GATTCACCAA	ATGTGAGTGT	GAGCTGTTTC	ATTCCAAAGT	AGGACTGGAC '	TATCTTTTTC	180
TCCTCAAAAC	TGGAAACCAA	AACGGCATCC	AAAGGACCTT	TTGACCTCGC	CAGGAGAAAG	240
TCATGCATAT	TATGGAAGAT	TGCATAATCA	GCGAAAGCGG	CAGTCATTGA	GCCCTTTTTG	300
AATTGACATG	GCAGTCTTGA	TTCGAACATT	GGATTCACAA	ACCTCCTGCT	TTCAATGTTA	360
TACCTATACG	TCTTGCCCTC	AAGCAGTCTA	AGTATGTCAC	TCTGCTCAGC	GCCCAAAATT	420
GCCCGATCCT	TTTGCGAATC	ATATGTGCTT	TGACATGGAT	CACCGGCAAG	TGAAATGAAA	480
GCATCACTAC	GTATAATAAG	CAAGCATAGA	TCGATGTATC	CAGGAGGAAA	CAACTGGACC	540
TCATCGAAAA	TCACACAGTG	GCCAGCCTTC	AATCCTGCAA	CTTTTCTGAG	GAAAACCTCA	600
AAAGTTACTA	CATCCCACCC	TTCCTTCTTT	GACCTACCTG	CTTTAGCAAC	TTTGCAGCTA	660
TCATCCATTT	CAAGATCATT	TTTGATTGAA	TTTGCTAAAG	CACGTCTGGG	AGAAACAAAG	720
GTTACGAATT	TACCCTCAGA	CCTTTTCATG	AAACTCTTGT	ACAAGAAACT	CTTCCCAGCC	780
CCAAATGTAC	CAAGCACGAC	AGTCAACTCC	CTTGGCTTAA	TATCAGTAGT	AGATATACCA	840
GAAAGCCAAG	GTTTTGCATC	ACTGAACTTC	TCATCACTTA	TAACGCCAGT	TAGGCCCCCT	900
AGCAAAC					`	907
	7 79 1	.• •	C140.04.50	(TDC + D 1)	1	
ID. No. 40 a		uae sequence	01 140-94-72	(1/+R1) corre	esponds to SEQ.	
1D. No. 40 a	5 10110WS.					
AGAATGCTTA	TGCTGAGAAT	GAGATGATT	CATTATTTTG	CATCCGGCAC	CATGTAAGGC	60
TTATAGTAAT	' AACACCGGAA	TATGAAGTT	A GTTGGAAATT	TGGGGAAAGT	GAGTGGCCCC	120
TATGTGGAAT	TCTTTGCCTG	AGGTCCAATO	CACTTCCAACC	ATGCGCCCCG	CTGAATGGTT	180
GCATGATCAC	GGCTATTGCT	TCAGCACTTO	G GGAGGCGTGA	GGTTGATGTG	TTAAATTATC	240
TGTGTAGGCC	TAGCACTAAT	CACATCTTT	G AGGAGCTGTG	CCAGGGCGGA	GGGCTTAATA	300
TGATGTACTI	GGCTGAAGCT	TTTGAGGCC	r TTGACATTTC	TGCAAAGTGC	GACATAAATG	360

ATGACTGCTG CCTTTGCTGA TTATGCAATC TT

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GGGAAATT	σA	GGTCATTAAC	CCAAATGGCA	AGATTTCCGC	CIIGIIIGAI	ATAACTAATG	420
AGCACATA	AG	GCATGTTGAG	AAGATCAGCA	ATGGCCCTCA	GAGCATAAAA	ATAGATGAGT	480
TGAGGAAG	GT	TAAGCGATCC	CGCCTTGACC	TTCTTTCAAT	GAATGGGTCC	AAAATAACCT	540
ATTTTCCA	AA	CTTTGAGCGG	GCTGAAAAGT	TGCAAGGGTG	CTTGCTAGAG	GGCĆTGACTG	600
GTGTCATA	AG	TGATGAAAAG	TTCAGTGATG	CAAAACCTTG	GCTTTCTGGT	ATATCAACTG	660
CGGATATT	AA	GCCAAGAGAG	CTAACTGTCG	TGCTTGGCAC	ATTTGGTGCT	GGAAAGAGTT	720
TCTTGTAT	ΑA	GAGTTTCATG	AAGAGATCTG	AAGGAAAATT	TGTAACTTTT	GTTTCCCCTA	780
GGCGAGCT	TT	GGCCAATTCG	ATCAAGAATG	ATCTTGAAAT	GGATGATGGC	TGCAAAGTTG	840
CCAAAGCA	.GG	CAAGTCAAAG	AAGGAAGGGT	GGGATGTGGT	AACATTTGAG	GTTTTCCTTA	900
GAAAAGTT	TC	TGGTTTGAAG	GCTGGTCATT	GTGTGATTTT	CGATGAGGTT	CAGTTGTTTC	960
CCCCTGGA	TA.	TATCGATCTA	TGTTTACTTG	TCATACGCAG	TGATGCTTTT	ATTTCACTTG	1020

CCGGTGATCC ATGCCAGAGC ACATATGATT CACAAAAGGA TCGGGCAATT TTGGGAGCTG

AGCAGAGTGA CATACTCAGA TTGCTTGAAG GAAAGACGTA TAGGTACAAC ATAGAAAGCA

GACGTTTTGT GAACCCAATG TTTGAATTTA GACTACCATG TCACTTCAAA AAAGGGTTCA

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Also encompassed by the present invention are fragments of the DNA molecules of the present invention. Suitable fragments capable of imparting RSP resistance to grape plants are constructed by using appropriate restriction sites, revealed by inspection of the DNA molecule's sequence, to: (i) insert an interposon (Felley et al., "Interposon Mutagenesis of Soil and Water Bacteria: A Family of DNA Fragments Designed for in vitro Insertion Mutagenesis of Gram-negative Bacteria," Gene, 52:147-15 (1987), which is hereby incorporated by reference) such that truncated forms of the RSP virus polypeptide or protein, that lack various amounts of the C-terminus, can be produced or (ii) delete various internal portions of the protein. Alternatively, the sequence can be used to amplify any portion of the coding region, such that it can be cloned into a vector supplying both transcription and translation start signals.

Suitable DNA molecules are those that hybridize to a DNA molecule comprising a nucleotide sequence of at least 15 continuous bases of SEQ. ID. No. 1 under stringent conditions characterized by a hybridization buffer comprising 0.9M sodium citrate ("SSC") buffer at a temperature of 37°C and remaining bound when

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subject to washing with SSC buffer at 37°C; and preferably in a hybridization buffer comprising 20% formamide in 0.9M saline/0.9M SSC buffer at a temperature of 42°C and remaining bound when subject to washing at 42°C with 0.2x SSC buffer at 42°C.

Variants may also (or alternatively) be modified by, for example, the deletion or addition of nucleotides that have minimal influence on the properties, secondary structure and hydropathic nature of the encoded protein or polypeptide. For example, the nucleotides encoding a protein or polypeptide may be conjugated to a signal (or leader) sequence at the N-terminal end of the protein which cotranslationally or post-translationally directs transfer of the protein.—The nucleotide sequence may also be altered so that the encoded protein or polypeptide is conjugated to a linker or other sequence for ease of synthesis, purification, or identification of the polypeptide.

The protein or polypeptide of the present invention is preferably produced in purified form (preferably, at least about 80%, more preferably 90%, pure) by conventional techniques. Typically, the protein or polypeptide of the present invention is isolated by lysing and sonication. After washing, the lysate pellet is resuspended in buffer containing Tris-HCl. During dialysis, a precipitate forms from this protein solution. The solution is centrifuged, and the pellet is washed and resuspended in the buffer containing Tris-HCl. Proteins are resolved by electrophoresis through an SDS 12% polyacrylamide gel.

The DNA molecule encoding the RSP virus protein or polypeptide of the present invention can be incorporated in cells using conventional recombinant DNA technology. Generally, this involves inserting the DNA molecule into an expression system to which the DNA molecule is heterologous (i.e., not normally present). The heterologous DNA molecule is inserted into the expression system or vector in proper sense orientation and correct reading frame. The vector contains the necessary elements for the transcription and translation of the inserted protein-coding sequences.

U.S. Patent No. 4,237,224 to Cohen and Boyer, which is hereby
incorporated by reference, describes the production of expression systems in the form
of recombinant plasmids using restriction enzyme cleavage and ligation with DNA
ligase. These recombinant plasmids are then introduced by means of transformation

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and replicated in unicellular cultures including procaryotic organisms and eukaryotic cells grown in tissue culture.

Recombinant genes may also be introduced into viruses, such as vaccinia virus. Recombinant viruses can be generated by transfection of plasmids into cells infected with virus.

Suitable vectors include, but are not limited to, the following viral vectors such as lambda vector system gt11, gt WES.tB, Charon 4, and plasmid vectors such as pBR322, pBR325, pACYC177, pACYC184, pUC8, pUC9, pUC18, pUC19, pLG339; pR290, pKC37, pKC101, SV 40, pBluescript II SK +/- or KS +/- (see "Stratagene Cloning Systems" Catalog (1993) from Stratagene, La Jolla, Calif, which is hereby incorporated by reference), pQE, pIH821, pGEX, pET series (see Studier et. al., "Use of T7 RNA Polymerase to Direct Expression of Cloned Genes," Gene Expression Technology, vol. 185 (1990), which is hereby incorporated by reference), and any derivatives thereof. Suitable vectors are continually being developed and identified. Recombinant molecules can be introduced into cells via transformation, transduction, conjugation, mobilization, or electroporation. The DNA sequences are cloned into the vector using standard cloning procedures in the art, as described by Maniatis et al., Molecular Cloning: A Laboratory Manual, Cold Springs Laboratory, Cold Springs Harbor, New York (1982), which is hereby incorporated by reference.

A variety of host-vector systems may be utilized to express the proteinencoding sequence(s). Primarily, the vector system must be compatible with the host cell used. Host-vector systems include but are not limited to the following: bacteria transformed with bacteriophage DNA, plasmid DNA, or cosmid DNA; microorganisms such as yeast containing yeast vectors; mammalian cell systems infected with virus (e.g., vaccinia virus, adenovirus, etc.); insect cell systems infected with virus (e.g., baculovirus); and plant cells infected by bacteria or transformed via particle bombardment (i.e. biolistics). The expression elements of these vectors vary in their strength and specificities. Depending upon the host-vector system utilized, any one of a number of suitable transcription and translation elements can be used.

Different genetic signals and processing events control many levels of gene expression (e.g., DNA transcription and messenger RNA ("mRNA") translation).

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Transcription of DNA is dependent upon the presence of a promoter which is a DNA sequence that directs the binding of RNA polymerase and thereby promotes mRNA synthesis. The DNA sequences of eukaryotic promoters differ from those of procaryotic promoters. Furthermore, eukaryotic promoters and accompanying genetic signals may not be recognized in or may not function in a procaryotic system, and, further, procaryotic promoters are not recognized and do not function in eukaryotic cells.

Similarly, translation of mRNA in procaryotes depends upon the presence of the proper procaryotic signals which differ from those of eukaryotes. Efficient translation of mRNA in procaryotes requires a ribosome binding site called the Shine-Dalgarno ("SD") sequence on the mRNA. This sequence is a short nucleotide sequence of mRNA that is located before the start codon, usually AUG, which encodes the amino-terminal methionine of the protein. The SD sequences are complementary to the 3'-end of the 16S rRNA (ribosomal RNA) and probably promote binding of mRNA to ribosomes by duplexing with the rRNA to allow correct positioning of the ribosome. For a review on maximizing gene expression, see Roberts and Lauer, Methods in Enzymology, 68:473 (1979), which is hereby incorporated by reference.

Promoters vary in their "strength" (i.e., their ability to promote transcription). For the purposes of expressing a cloned gene, it is desirable to use strong promoters in order to obtain a high level of transcription and, hence, expression of the gene. Depending upon the host cell system utilized, any one of a number of suitable promoters may be used. For instance, when cloning in *E. coli*, its bacteriophages, or plasmids, promoters such as the T7 phage promoter, *lac* promoter, *trp* promoter, *rec*A promoter, ribosomal RNA promoter, the P_R and P_L promoters of coliphage lambda and others, including but not limited, to *lac*UV5, *omp*F, *bla*, *lpp*, and the like, may be used to direct high levels of transcription of adjacent DNA segments. Additionally, a hybrid *trp-lac*UV5 (*tac*) promoter or other *E. coli* promoters produced by recombinant DNA or other synthetic DNA techniques may be used to provide for transcription of the inserted gene.

Bacterial host cell strains and expression vectors may be chosen which inhibit the action of the promoter unless specifically induced. In certain operons, the addition of specific inducers is necessary for efficient transcription of the inserted

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DNA. For example, the *lac* operon is induced by the addition of lactose or IPTG (isopropylthio-beta-D-galactoside). A variety of other operons, such as *trp*, *pro*, etc.,

are under different controls.

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Specific initiation signals are also required for efficient gene transcription and translation in procaryotic cells. These transcription and translation initiation signals may vary in "strength" as measured by the quantity of gene specific messenger RNA and protein synthesized, respectively. The DNA expression vector, which contains a promoter, may also contain any combination of various "strong" transcription and/or translation initiation signals. For instance, efficient translation in *E. coli* requires a Shine-Dalgarno ("SD") sequence about 7-9 bases 5' to the initiation codon ("ATG") to provide a ribosome binding site. Thus, any SD-ATG combination that can be utilized by host cell ribosomes may be employed. Such combinations include but are not limited to the SD-ATG combination from the *cro* gene or the *N* gene of coliphage lambda, or from the *E. coli* tryptophan E, D, C, B or A genes. Additionally, any SD-ATG combination produced by recombinant DNA or other techniques involving incorporation of synthetic nucleotides may be used.

Once the isolated DNA molecules encoding the various *Rupestris* stem pitting associated virus proteins or polypeptides, as described above, have been cloned into an expression system, they are ready to be incorporated into a host cell. Such incorporation can be carried out by the various forms of transformation noted above, depending upon the vector/host cell system. Suitable host cells include, but are not limited to, bacteria, virus, yeast, mammalian cells, insect, plant, and the like.

The present invention also relates to RNA molecules which encode the various RSP virus proteins or polypeptides described above. The transcripts can be synthesized using the host cells of the present invention by any of the conventional techniques. The mRNA can be translated either *in vitro* or *in vivo*. Cell-free systems typically include wheat-germ or reticulocyte extracts. *In vivo* translation can be effected, for example, by microinjection into frog oocytes.

One aspect of the present invention involves using one or more of the above DNA molecules encoding the various proteins or polypeptides of a RSP virus to transform grape plants in order to impart RSP resistance to the plants. The mechanism by which resistance is imparted in not known. In one hypothetical mechanism, the transformed plant can express the coat protein or polypeptide, and,

when the transformed plant is inoculated by a RSP virus, such as RSPaV-1, the expressed coat protein or polypeptide surrounds the virus, thereby preventing translation of the viral DNA.

In this aspect of the present invention, the subject DNA molecule incorporated in the plant can be constitutively expressed. Alternatively, expression can be regulated by a promoter which is activated by the presence of RSP virus. Suitable promoters for these purposes include those from genes expressed in response to RSP virus infiltration.

The isolated DNA molecules of the present invention can be utilized to impart RSP virus resistance for a wide variety of grapevine plants. The DNA 10 molecules are particularly well suited to imparting resistance to Vitis scion or rootstock cultivars. Scion cultivars which can be protected include those commonly referred to as Table or Raisin Grapes, such as Alden, Almeria, Anab-E-Shahi, Autumn Black, Beauty Seedless, Black Corinth, Black Damascus, Black Malvoisie, Black Prince, Blackrose, Bronx Seedless, Burgrave, Calmeria, Campbell Early, 15 Canner, Cardinal, Catawba, Christmas, Concord, Dattier, Delight, Diamond, Dizmar, Duchess, Early Muscat, Emerald Seedless, Emperor, Exotic, Ferdinand de Lesseps, Fiesta, Flame seedless, Flame Tokay, Gasconade, Gold, Himrod, Hunisa, Hussiene, Isabella, Italia, July Muscat, Khandahar, Katta, Kourgane, Kishmishi, Loose Perlette, Malaga, Monukka, Muscat of Alexandria, Muscat Flame, Muscat Hamburg, New 20 York Muscat, Niabell, Niagara, Olivette blanche, Ontario, Pierce, Queen, Red Malaga, Ribier, Rish Baba, Romulus, Ruby Seedless, Schuyler, Seneca, Suavis (IP 365), Thompson seedless, and Thomuscat. They also include those used in wine production, such as Aleatico, Alicante Bouschet, Aligote, Alvarelhao, Aramon, Baco blanc (22A), Burger, Cabernet franc, Cabernet, Sauvignon, Calzin, Carignane, 25 Charbono, Chardonnay, Chasselas dore, Chenin blanc, Clairette blanche, Early Burgundy, Emerald Riesling, Feher Szagos, Fernao Pires, Flora, French Colombard, Fresia, Furmint, Gamay, Gewurztraminer, Grand noir, Gray Riesling, Green Hungarian, Green Veltliner, Grenache, Grillo, Helena, Inzolia, Lagrein, Lambrusco de Salamino, Malbec, Malvasia bianca, Mataro, Melon, Merlot, Meunier, Mission, 30 Montua de Pilas, Muscadelle du Bordelais, Muscat blanc, Muscat Ottonel, Muscat Saint-Vallier, Nebbiolo, Nebbiolo fino, Nebbiolo Lampia, Orange Muscat, Palomino, Pedro Ximenes, Petit Bouschet, Petite Sirah, Peverella, Pinot noir, Pinot Saint-

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George, Primitivo di Gioa, Red Veltliner, Refosco, Rkatsiteli, Royalty, Rubired, Ruby Cabernet, Saint-Emilion, Saint Macaire, Salvador, Sangiovese, Sauvignon blanc, Sauvignon gris, Sauvignon vert, Scarlet, Seibel 5279, Seibel 9110, Seibel 13053, Semillon, Servant, Shiraz, Souzao, Sultana Crimson, Sylvaner, Tannat, Teroldico,
Tinta Madeira, Tinto cao, Touriga, Traminer, Trebbiano Toscano, Trousseau, Valdepenas, Viognier, Walschriesling, White Riesling, and Zinfandel. Rootstock cultivars which can be protected include Couderc 1202, Couderc 1613, Couderc 1616, Couderc 3309, Dog Ridge, Foex 33 EM, Freedom, Ganzin 1 (A x R #1), Harmony, Kober 5BB, LN33, Millardet & de Grasset 41B, Millardet & de Grasset 420A,
Millardet & de Grasset 101-14, Oppenheim 4 (SO4), Paulsen 775, Paulsen 1045, Paulsen 1103, Richter 99, Richter 110, Riparia Gloire, Ruggeri 225, Saint-George, Salt Creek, Teleki 5A, Vitis rupestris Constantia, Vitis california, and Vitis girdiana.

Plant tissue suitable for transformation include leaf tissue, root tissue, meristems, zygotic and somatic embryos, and anthers. It is particularly preferred to utilize embryos obtained from anther cultures.

The expression system of the present invention can be used to transform virtually any plant tissue under suitable conditions. Tissue cells transformed in accordance with the present invention can be grown *in vitro* in a suitable medium to impart RSPaV resistance. Transformed cells can be regenerated into whole plants such that the protein or polypeptide imparts resistance to RSPaV in the intact transgenic plants. In either case, the plant cells transformed with the recombinant DNA expression system of the present invention are grown and caused to express that DNA molecule to produce one of the above-described RSPaV proteins or polypeptides and, thus, to impart RSPaV resistance.

In producing transgenic plants, the DNA construct in a vector described above can be microinjected directly into plant cells by use of micropipettes to transfer mechanically the recombinant DNA. Crossway, Mol. Gen. Genetics, 202:179-85 (1985), which is hereby incorporated by reference. The genetic material may also be transferred into the plant cell using polyethylene glycol. Krens, et al., Nature, 296:72-74 (1982), which is hereby incorporated by reference.

One technique of transforming plants with the DNA molecules in accordance with the present invention is by contacting the tissue of such plants

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with an inoculum of a bacteria transformed with a vector comprising a gene in accordance with the present invention which imparts RSPaV resistance.

Generally, this procedure involves inoculating the plant tissue with a suspension of bacteria and incubating the tissue for 48 to 72 hours on regeneration medium without antibiotics at 25-28°C.

Bacteria from the genus Agrobacterium can be utilized to transform plant cells. Suitable species of such bacterium include Agrobacterium tumefaciens and Agrobacterium rhizogenes. Agrobacterium tumefaciens (e.g., strains C58, LBA4404, or EHA105) is particularly useful due to its well-known ability to transform plants.

Heterologous genetic sequences can be introduced into appropriate plant cells, by means of the Ti plasmid of A. tumefaciens or the Ri plasmid of A. rhizogenes. The Ti or Ri plasmid is transmitted to plant cells on infection by Agrobacterium and is stably integrated into the plant genome. J. Schell, Science, 237:1176-83 (1987), which is hereby incorporated by reference.

After transformation, the transformed plant cells must be regenerated.

Plant regeneration from cultured protoplasts is described in Evans et al., Handbook of Plant Cell Cultures, Vol. 1: (MacMillan Publishing Co., New York, 1983); and Vasil I.R. (ed.), Cell Culture and Somatic Cell Genetics of Plants, Acad. Press, Orlando, Vol. I, 1984, and Vol. III (1986), which are hereby incorporated by reference.

It is known that practically all plants can be regenerated from cultured cells or tissues, including but not limited to, all major species of sugarcane, sugar beets, cotton, fruit trees, and legumes.

Means for regeneration vary from species to species of plants, but generally a suspension of transformed protoplasts or a petri plate containing explants is first provided. Callus tissue is formed and shoots may be induced from callus and subsequently rooted. Alternatively, embryo formation can be induced in the callus tissue. These embryos germinate as natural embryos to form plants. The culture media will generally contain various amino acids and hormones, such as auxin and cytokinins. It is also advantageous to add glutamic acid and proline to the medium. Efficient regeneration will depend on the medium, on the genotype, and on the history

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of the culture. If these three variables are controlled, then regeneration is usually reproducible and repeatable.

After the expression cassette is stably incorporated in transgenic plants, it can be transferred to other plants by sexual crossing. Any of a number of standard breeding techniques can be used, depending upon the species to be crossed.

Once transgenic plants of this type are produced, the plants themselves can be cultivated in accordance with conventional procedure so that the DNA construct is present in the resulting plants. Alternatively, transgenic seeds are recovered from the transgenic plants. These seeds can then be planted in the soil and cultivated using conventional procedures to produce transgenic plants.

Another approach to transforming plant cells with a gene which imparts resistance to pathogens is particle bombardment (also known as biolistic transformation) of the host cell. This can be accomplished in one of several ways. The first involves propelling inert or biologically active particles at cells. This technique is disclosed in U.S. Patent Nos. 4,945,050, 5,036,006, and 5,100,792, all to Sanford et al., and in Emerschad et al., "Somatic Embryogenesis and Plant Development from Immature Zygotic Embryos of Seedless Grapes (Vitis vinifera)," Plant Cell Reports, 14:6-12 (1995) ("Emerschad (1995)"), which are hereby incorporated by reference. Generally, this procedure involves propelling inert or biologically active particles at the cells under conditions effective to penetrate the outer surface of the cell and to be incorporated within the interior thereof. When inert particles are utilized, the vector can be introduced into the cell by coating the particles with the vector containing the heterologous DNA. Alternatively, the target cell can be surrounded by the vector so that the vector is carried into the cell by the wake of the particle. Biologically active particles (e.g., dried bacterial cells containing the vector and heterologous DNA) can also be propelled into plant cells.

Once a grape plant tissue is transformed in accordance with the present invention, it is regenerated to form a transgenic grape plant. Generally, regeneration is accomplished by culturing transformed tissue on medium containing the appropriate growth regulators and nutrients to allow for the initiation of shoot meristems. Appropriate antibiotics are added to the regeneration medium to inhibit the growth of Agrobacterium and to select for the

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development of transformed cells. Following shoot initiation, shoots are allowed to develop tissue culture and are screened for marker gene activity.

The DNA molecules of the present invention can be made capable of transcription to a messenger RNA that does not translate to the protein. This is known as RNA-mediated resistance. When a *Vitis* scion or rootstock cultivar is transformed with such a DNA molecule, the DNA molecule can be transcribed under conditions effective to maintain the messenger RNA in the plant cell at low level density readings. Density readings of between 15 and 50 using a Hewlet ScanJet and Image Analysis Program are preferred.

A portion of one or more DNA molecules of the present invention as well as other DNA molecules can be used in a transgenic grape plant in accordance with U.S. Patent Application Serial No. 09/025,635, which is hereby incorporated herein by reference.

The RSPaV protein or polypeptide can also be used to raise antibodies or binding portions thereof or probes. The antibodies can be monoclonal or polyclonal.

Monoclonal antibody production may be effected by techniques which are well-known in the art. Basically, the process involves first obtaining immune cells (lymphocytes) from the spleen of a mammal (e.g., mouse) which has been previously immunized with the antigen of interest either *in vivo* or *in vitro*. The antibody-secreting lymphocytes are then fused with (mouse) myeloma cells or transformed cells, which are capable of replicating indefinitely in cell culture, thereby producing an immortal, immunoglobulin-secreting cell line. The resulting fused cells, or hybridomas, are cultured, and the resulting colonies screened for the production of the desired monoclonal antibodies. Colonies producing such antibodies are cloned, and grown either *in vivo* or *in vitro* to produce large quantities of antibody. A description of the theoretical basis and practical methodology of fusing such cells is set forth in Kohler and Milstein, Nature, 256:495 (1975), which is hereby incorporated by reference.

Mammalian lymphocytes are immunized by *in vivo* immunization of the animal (e.g., a mouse) with the protein or polypeptide of the present invention. Such immunizations are repeated as necessary at intervals of up to

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several weeks to obtain a sufficient titer of antibodies. Following the last antigen boost, the animals are sacrificed and spleen cells removed.

Fusion with mammalian myeloma cells or other fusion partners capable of replicating indefinitely in cell culture is effected by standard and well-known techniques, for example, by using polyethylene glycol ("PEG") or other fusing agents. (See Milstein and Kohler, Eur. J. Immunol., 6:511 (1976), which is hereby incorporated by reference.) This immortal cell line, which is preferably murine, but may also be derived from cells of other mammalian species, including but not limited to rats and humans, is selected to be deficient in enzymes necessary for the utilization of certain nutrients, to be capable of rapid growth, and to have good fusion capability. Many such cell lines are known to those skilled in the art, and others are regularly described.

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Procedures for raising polyclonal antibodies are also well known. Typically, such antibodies can be raised by administering the protein or polypeptide of the present invention subcutaneously to New Zealand white rabbits which have first been bled to obtain pre-immune serum. The antigens can be injected at a total volume of 100 µl per site at six different sites. Each injected material will contain synthetic surfactant adjuvant pluronic polyols, or pulverized acrylamide gel containing the protein or polypeptide after SDS-polyacrylamide gel electrophoresis. The rabbits are then bled two weeks after the first injection and periodically boosted with the same antigen three times every six weeks. A sample of serum is then collected 10 days after each boost. Polyclonal antibodies are then recovered from the serum by affinity chromatography using the corresponding antigen to capture the antibody. Ultimately, the rabbits are euthanized with pentobarbital 150 mg/Kg IV. This and other procedures for raising polyclonal antibodies are disclosed in Harlow et. al., editors. Antibodies: A Laboratory Manual (1988), which is hereby incorporated by reference.

In addition to utilizing whole antibodies, binding portions of such antibodies can be used. Such binding portions include Fab fragments, F(ab')₂ fragments, and Fv fragments. These antibody fragments can be made by conventional procedures, such as proteolytic fragmentation procedures, as

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described in Goding, Monoclonal Antibodies: Principles and Practice, New York: Academic Press, pp. 98-118 (1983), which is hereby incorporated by reference.

The present invention also relates to probes found either in nature or prepared synthetically by recombinant DNA procedures or other biological procedures. Suitable probes are molecules that bind to RSP viral antigens identified by the polyclonal antibodies of the present invention or bind to the nucleic acid of RSPaV. Such probes can be, for example, proteins, peptides, lectins, or nucleic acids.

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The antibodies or binding portions thereof or probes can be
administered to RSPaV infected scion cultivars or rootstock cultivars.

Alternatively, at least the binding portions of these antibodies can be sequenced, and the encoding DNA synthesized. The encoding DNA molecule can be used to transform plants together with a promoter which causes expression of the encoded antibody when the plant is infected by an RSPaV. In either case, the antibody or binding portion thereof or probe will bind to the virus and help prevent the usual stem pitting response.

Antibodies raised against the proteins or polypeptides of the present invention or binding portions of these antibodies can be utilized in a method for detection of RSPaV in a sample of tissue, such as tissue from a grape scion or rootstock. Antibodies or binding portions thereof suitable for use in the detection method include those raised against a replicase, proteins or polypeptides of the triple gene block, or a coat protein or polypeptide in accordance with the present invention. Any reaction of the sample with the antibody is detected using an assay system which indicates the presence of RSPaV in the sample. A variety of assay systems can be employed, such as enzyme-linked immunosorbent assays, radioimmunoassays, gel diffusion precipitin reaction assays, immunodiffusion assays, agglutination assays, fluorescent immunoassays, protein A immunoassays, or immunoelectrophoresis assays.

Alternatively, the RSPaV can be detected in such a sample using the

30 DNA molecules of the present, RNA molecules of the present invention, or DNA or
RNA fragments thereof, as probes in nucleic acid hybridization assays for detecting
the presence of complementary virus DNA or RNA in the various tissue samples
described above. The nucleotide sequence is provided as a probe in a nucleic acid

hybridization assay or a gene amplification detection procedure (e.g., using a polymerase chain reaction procedure). The nucleic acid probes of the present invention may be used in any nucleic acid hybridization assay system known in the art, including, but not limited to, Southern blots (Southern, E.M., "Detection of Specific Sequences Among DNA Fragments Separated by Gel Electrophoresis," 5 J. Mol. Biol., 98:503-17 (1975), which is hereby incorporated by reference), Northern blots (Thomas, P.S., "Hybridization of Denatured RNA and Small DNA Fragments Transferred to Nitrocellulose," Proc. Nat'l Acad. Sci. USA, 77:5201-05 (1980), which is hereby incorporated by reference), and Colony blots (Grunstein, M., et al., "Colony 10 Hybridization: A Method for the Isolation of Cloned cDNAs that Contain a Specific Gene," Proc. Nat'l Acad. Sci. USA, 72:3961-65 (1975), which is hereby incorporated by reference). Alternatively, the isolated DNA molecules of the present invention or RNA transcripts thereof can be used in a gene amplification detection procedure (e.g., a polymerase chain reaction). Erlich, H.A., et. al., "Recent Advances in the Polymerase Chain Reaction," Science 252:1643-51 (1991), which is hereby 15 incorporated by reference. Any reaction with the probe is detected so that the presence of RSP virus in the sample is indicated. Such detection is facilitated by providing the DNA molecule of the present invention with a label. Suitable labels include a radioactive compound, a fluorescent compound, a chemiluminescent compound, an enzymatic compound, or other equivalent nucleic acid labels. 20

Depending upon the desired scope of detection, it is possible to utilize probes having nucleotide sequences that correspond with conserved or variable regions of the ORF or UTR. For example, to distinguish RSPaV from other related viruses (as described herein), it is desirable to use probes which contain nucleotide sequences that correspond to sequences more highly conserved among all RSPaV strains. Also, to distinguish between different RSPaV strains (e.g., RSPaV-1, RSP47-4, RSP158), it is desirable to utilize probes containing nucleotide sequences that correspond to sequences less highly conserved among the RSP virus strains.

Nucleic acid (DNA or RNA) probes of the present invention will

hybridize to complementary RSPaV-1 nucleic acid under stringent conditions. Less stringent conditions may also be selected. Generally, stringent conditions are selected to be about 50°C lower than the thermal melting point (T_m) for the specific sequence at a defined ionic strength and pH. The T_m is the temperature (under defined ionic

strength and pH) at which 50% of the target sequence hybridizes to a perfectly matched probe. The T_m is dependent upon the solution conditions and the base composition of the probe, and may be calculated using the following equation:

$$T_m = 79.8^{\circ}C + (18.5 \times Log[Na+])$$
+ $(58.4^{\circ}C \times \%[G+C])$
- $(820 / \#bp \text{ in duplex})$
- $(0.5 \times \% \text{ formamide})$

Nonspecific binding may also be controlled using any one of a number of known techniques such as, for example, blocking the membrane with protein-containing solutions, addition of heterologous RNA, DNA, and SDS to the hybridization buffer, and treatment with RNase. Generally, suitable stringent conditions for nucleic acid hybridization assays or gene amplification detection procedures are as set forth above. More or less stringent conditions may also be selected.

The development of a rapid detection method for RSP is a major breakthrough, because the only detection method now available is through inoculation of St. George grape indicators, which takes two to three years to develop symptoms. A serological or nucleic acid based detection tests developed for RSP will take only 1 to 2 days and it is less expensive. The woody indicator test on St. George costs \$250 per sample, while a serological or nucleic acid based test would cost \$30-50 per sample. Moreover, the rapid tests will speed up the introduction of grape imports into the US from the current three years to about six months. These applications will be valuable wherever grapes are grown. Since RSP is part of the rugose wood complex, development of rapid detection methods will be invaluable in determining the significance of RSP in the rugose wood complex. This will allow an investigator to determine whether RSP alone can cause the rugose wood complex or if other components are needed. In addition, these rapid detection methods are very useful to evaluate the resistance of transgenic plants to Rupestris stem pitting associated virus.

EXAMPLES

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The following examples are provided to illustrate embodiments of the present invention but are by no means intended to limit its scope.

Example 1 - Grapevine Materials for dsRNA Analysis

Samples from 15 accessions that induced pitting on graft-inoculated
St. George were collected from the National Grapevine Germplasm Repository of the
USDA Plant Genetic Resources Unit (PGRU) at Geneva and used for dsRNA
analysis. Positive controls used included Thompson Seedless (RSP105) (Golino,
"The Davis Grapevine Virus Collection," Am. J. Enology Viticulture, 43:200-05
(1992), which is hereby incorporated by reference) from the FPMS, University of
California (Davis) and Pinot Noir (SVP1186-09A2), which was kindly provided by
Dr. R. Johnson of Center for Plant Health, Agriculture Canada, Sidney, British
Columbia. Negative controls as judged by indexing on St. George included Freedom
from the PGRU at Geneva, New York, and Verduzzo 233A. The latter was kindly
provided by Dr. P. Silvano of the Sezione di Fitovirologia, ERSA Servizio ChimicoAgrario e della Certificazione, Pozzuolo del Friuh (UD), Italy.

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Example 2 - Grapevine Materials for RT-PCR

Dormant cuttings of 138 grapevine selections were collected from USA, Canada, Italy, and Portugal over three years. Samples included *Vitis* vinifera cultivars, hybrids, V. riparia, and rootstocks. 117 grapevine selections were indexed on St. George for RSP and other RW diseases. Pinot noir (1186-9A2) from Agriculture Canada, Center for Plant Health (Sidney, Canada) and Thompson seedless (RSP105) from University of California (Davis) were included as positive controls. Sauvignon blanc, generated from shoot tip tissue culture and tested free of viruses and viroids was provided by Dr. J. Semancik (University of California at Riverside) and used as a healthy control. In addition, six seedlings of five *Vitis* species were also included as negative controls.

Example 3 - dsRNA Isolation and Analysis

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Methods for isolating dsRNA were described by Hu et al., "Characterization of Closterovirus-like Particles Associated with Grapevine Leafroll Disease," J. Phytopathology, 128:1-14 (1990), which is hereby incorporated by

reference, except that 1 X STE with 15% ethanol (instead of 16.5%) was used to wash CF-11 cellulose columns prior to elution of dsRNAs. The dsRNAs were isolated from leaves, petioles, and the phloem tissue of dormant canes, electrophoresed on 1% agarose or low melting temperature agarose gels, and analyzed by staining with ethidium bromide (EtBr). *Hind* EII digested lambda DNA was used as markers to estimate the sizes of the dsRNA molecules.

Example 4 - cDNA Synthesis and Cloning

The extremely low yield of dsRNA and the limited quantity of RSP-10 infected grape materials precluded the use of a single RSP-infected grapevine accession as the source of dsRNA for cloning purpose. Therefore, dsRNA preparations from Colobel 257, Ravat 34, Couderc 28-112, and Seyval were pooled and used as templates for cDNA synthesis. In order to get pure templates for cloning, dsRNA bands were excised from low melting temperature agarose gels after 15 electrophoresis and recovered by extraction with phenol and chloroform (Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY (1989), which is hereby incorporated by reference). The same recovery procedure was repeated once more. The purified dsRNA was denatured with 20 mM methyl mercuric hydroxide and cDNAs were 20 synthesized using slightly modified methods of Jelkmann et al., "Cloning of Four Viruses from Small Quantities of Double-Stranded RNA," Phytopathology, 79:1250-53 (1989), which is incorporated herein be reference. The cDNA fragments were first blunt-ended with T4 DNA polymerase at 12°C. T4 DNA ligase was used to add EcoR I adapters to both ends of the cDNAs. Subsequently, the cDNA molecules with 25 cohesive ends were ligated to EcoR I-prepared arms of lambda ZAP II. Finally, the resulting recombinant phages were packed into Gigapack II packaging extract following manufacturer's instructions (Stratagene, La Jolla, CA).

30 Example 5 - Identification of cDNA Clones Specific to the dsRNA

Plaque hybridization was used to screen cDNA clones by transferring recombinant cDNA plaques to nylon membranes and hybridizing to ³²P-labeled first-

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strand cDNA probes generated from the dsRNA according to manufacturer's recommendations (Du Pont, 1987). Clones with strong hybridization signals were converted into pBluescript SK through *in vivo* excision (Stratagene, 1991). After digestion of the resulting plasmids with *EcoR* I, 20 clones were selected and further analyzed in Southern hybridization with radio labeled first strand cDNA probes synthesized from the dsRNA. The specificity of two selected clones to the dsRNA was confirmed by Northern analysis using ³²P labeled inserts of the two clones.

Example 6 - Bridging Gaps Between Clones

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To bridge the gap between clones RSP3 and RSP94, a pair of specific primers were used in RT-PCR to generate cDNA fragments from the dsRNA. RSP3-RSP94 primer 1 (sense, nt 3629-3648) has a nucleotide sequence corresponding to SEO. ID. No. 41 as follows:

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GCTTCAGCAC TTGGAAGGCG

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RSP3-RSP94 primer 2 (antisense, nt 4350-4366) has a nucleotide sequence corresponding to SEQ. ID. No. 42 as follows:

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CACACAGTGG CCAGCCT

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After gel electrophoresis, PCR amplified cDNA bands were excised from gels and recovered with the phenol/chloroform method (Sambrook et al., Molecular Cloning:

A Laboratory Manual, 2nd ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY (1989), which is hereby incorporated by reference).

The same strategy was employed to bridge the gap between clones RSP94 and RSP95. RSP94-RSP95 primer 1 (sense, nt 5272-5291) has a nucleotide sequence corresponding to SEQ. ID. No. 43 as follows:

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GGAGGTGCGT TGTGGTTATG

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RSP94-RSP95 primer 2 (antisense, nt 6791-6808) has a nucleotide sequence corresponding to SEQ. ID. No. 44 as follows:

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CCCTGGCACT GCACACCC

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Example 7 - Obtaining Nucleotide Sequences on Both Termini of RSPaV-1 Genome

To obtain the terminal 3' end sequences, a primer (sense, nt 8193-8210) having a nucleotide sequence corresponding to SEQ. ID. No. 45 as follows:

GGAGGTGACC ACATTACG

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and a (dT)18 primer were used in RT-PCR to amplify cDNA from the dsRNA.

Resulting PCR products were cloned into TA vector pCRII (Invitrogen) and sequenced. This approach was based on the assumption that the RSP associated dsRNA contained a poly (A) tail. For the terminal 5' end, the dsRNA was first tagged with poly (A) using yeast Poly (A) polymerase (USB) (Pappu et al., "Nucleotide Sequence and Organization of Eight 3' Open Reading Frames of the Citrus tristeza

Closterovirus Genome," Virology 199:35-46 (1994), which is hereby incorporated by reference) and then used as templates to generate cDNA fragments by RT-PCR using

(dT)18 primer and primer (antisense, nt 429-449) having a nucleotide sequence

20 CATCACGACT TGTCACAAAC C

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Example 8 - Nucleotide Sequencing

corresponding to SEQ. ID. NO. 46 as follows:

CsCl or alkaline/PEG (polyethylene glycol) purified plasmids
(Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd ed., Cold Spring
Harbor Laboratory Press, Cold Spring Harbor, NY (1989), which is hereby
incorporated by reference; Applied Biosystems, Inc.) and RT-PCR amplified cDNA
fragments were sequenced for completion on both strands. Nucleotide sequencing
was done manually with Sequenase version 2.0 kit (USB) or automatically on ABI
373 automated sequencer with Taq DyeDeoxyTM terminator cycle sequencing kit
(Applied Biosystems, Inc.). Vector primers (T3, T7, M13 Forward, and M13
Reverse) were used in initial sequencing and sequences were completed by primer
walking strategy.

Example 9 - Reverse Transcription-Polymerase Chain Reaction (RT-PCR)

Two pairs of primers were designed for RT-PCR: (1) RSP95F1 and RSP95R1; and (2) RSP149F1 and RSP149R1. Primer RSP95F1, an antisense strand primer, has a nucleotide sequence corresponding to SEQ. ID. NO. 47 as follows: 5

TGGGCCTCCA CTTCTTC

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Primer RSP95R1, a sense strand primer, has a nucleotide sequence corresponding to SEO. ID. No. 48 as follows: 10

GGGGTTGCCT GAAGAT

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Primer RSP149F1, an antisense strand primer, has a nucleotide sequence corresponding to SEQ. ID. No. 49 as follows:

ACACCTGCTG TGAAAGC

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Primer RSP149R1, a sense strand primer, has a nucleotide sequence corresponding to SEQ. ID. No. 50 as follows:

GGCCAAGGTT CAGTTTG

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RSP95F1/R1 were used in RT-PCR to test samples collected in 1994. RSP149R1/F1, alone or together with RSP95F1/R1, were used to test samples collected in 1995 and 1996. To avoid bias in the judgment of RT-PCR results, blind tests were conducted for samples from Canada in 1995 and 1996. The indexing results of these samples were kept untold until the RT-PCR tests were complete.

dsRNAs were denatured with methylmercuric hydroxide (CH4HgOH) and reverse transcribed into cDNAs with Moloney murine leukemia virus (MMLV) or Avian Myeloblastosis Virus (AMV) reverse transcriptases (Promega) at 42 °C for 1 to 3 h. Five of 20 μ l of the RT reactions were added to PCR mix and amplified in thermal cycler (HYBAID OmniGene, National Labnet Company) with Taq DNA polymerase (buffer B, Promega) using the following parameters: initial denaturation at 94 °C for 5 min, 40 cycles of amplification at 94 °C for 45 s, 52 °C for 1 min, and 72 35 oC for 1 min, and a final extension at 72 oC for 10 min. PCR products were analyzed by electrophoresis on 1% agarose gels containing ethidium bromide. Hae III digested Phix 174 fragments were used as molecular weight markers.

Example 10 - Southern Blot

DNA fragments amplified by PCR from cDNA clone RSP149 with

primers RSP149F1/R1 were labeled with 32P by random priming and used as probes.

Products of RT-PCR of randomly selected grapevines including 26 positives and 6 negatives by RT-PCR were electrophoresed on an 0.8% agarose gel, transferred to nylon membranes, and hybridized to the probes following manufacturer's instructions (Du Pont).

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Example 11 - Computer Assisted Analysis of Sequences and Genome Structure of RSPaV-1

Sequences were assembled with SeqMan program and potential open 15 reading frames were generated with MapDraw program (DNASTAR, Madison, WI). BLAST program of the NCBI (the National Center for Biotechnology Information) was used to search for homologies in DNA and protein databases. Clustal analysis (with identity weight table) of MegAlign (DNASTAR) was employed to reveal sequence similarities between the putative proteins of RSPaV-1 and the analogous 20 proteins of ASPV (Jelkmann, "Nucleotide Sequences of Apple Stem Pitting Virus and of the Coat Protein of a Similar Virus from Pear Associated with Vein Yellows Disease and Their Relationship with Potex- and Carlaviruses," J. General Virology, 75:1535-42 (1994), which is hereby incorporated by reference) and PVM (Zavriev et al., "Complete Nucleotide Sequence of Genomic RNA of the Potato M-Virus." 25 Molecular Biology (Mosk.) 25:761-69 (1991), which is hereby incorporated by reference). In addition, nucleotide sequences of the untranslated regions (UTR) of these three viruses were also compared using MagAlign, as shown in Figures 6A and 6B.

30 Example 12 - Consistent Association of a High Molecular Weight dsRNA with RSP

The 15 grapevine accessions used in this study were previously indexed on St. George where 12 accessions induced typical RSP symptoms (i.e., a narrow strip of small pits below the inoculum bud). Figure 1A illustrates these

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typical RSP symptoms. A good correlation was found between the presence of the specific dsRNA and the indexing results on St. George. As shown in Figure 2A and recorded in Table 1 below, twelve grapevine accessions with typical RSP symptoms revealed a dsRNA of ca. 8.7 kb with gel electrophoresis. In addition, a smaller dsRNA of about 6.6 kb was observed in Colobel 257 and Seyval. In contrast, although Aminia and Canandaigua elicited deep pits and grooves around the woody cylinder of St. George, they did not reveal visible dsRNA of expected size in repeated experiments. Freedom, which indexed negative for RSP on St. George, did not reveal visible dsRNA. Although two dsRNA bands were observed in Verduzzo 233A (which was indexed free of RSP on St. George), they were not specific to RSP based on the fact that they were larger or smaller than the 8.7 kb dsRNA associated with RSP (Figure 2A) and that they did not hybridize to the RSP-specific probe in Northern analysis (Figure 2B). In addition, the two dsRNA species isolated from Verduzzo 233A were not observed in other healthy grapevines such as Cabernet Franc and LN 33.

Table 1

Accessions and Parentage	St. George Indicator	dsRNA	Northern
Aminia (Carter X Black Hamburg)	+	_	_
Bertille Seyve 3408 (BS 872 X Seibel 5410)	+	+	+
Bertille Seyve 5563 (Seibel 6905 X BS 3445)	+	+	+
Canandaigua (V. labrusca X V. vinifera)	+	_	<u>-</u> •
Colobel 257 (Seibel 6150 X Seibel 5455)	+ .	+	+
Couderc 28-112 (Emily X V. rupestris)	+ .	+	+
Freedom (Couderc 1613 X Dog Ridge)	_	· –	-
Grande Glabre (V. riparia)	+	+	+
III 344-1 (BS 2667 X Seibel 6905)	+	+†	_†
Joffre (V. vinifera X V. riparia X V. rupestris)	+	+	+
Ravat 34 (Berlandieri X Chardonnay)	+	+	+
Seyval (Seibel 4995 X Seibel 4986)	+	+ .	+
Seyve Villard 14-287 (V. labrusca X V. rupestris X V. aestiv X V. cinerea X V. vinifera)	+	+	+
Seyve Villard 3160 (Seibel 5163 X Seibel 2049)	+	+	+
Verdelet (Seibel 5455 X Seibel 4938)	+	+	+
Controls .			
Pinot Noir (V. vinifera)	+	+	+

Table 1

Accessions and Parentage	St. George Indicator	dsRNA	Northern
Thompson seedless (V. vinifera)	+	NT	+
Verduzzo 233A	-		_

Symbols:

The yield of dsRNA was low and varied significantly among different accessions. When a comparable amount of phloem tissue (14 g for Bertille Seyve 5563 and Couderc 28-112; 18.5 g for the others) was used to isolate dsRNA, Colobel 257, Seyval, Ravat 34, Grande Glabre, and Seyve Villard 14-287 displayed strong dsRNA bands, while Bertille Seyve 5563, Couderc 28-112, Joffre, and Verdelet showed weak bands after staining with EtBr, as shown in Figure 2A. Bertille Seyve 3408 and Seyve Villard 3160 were analyzed in separate experiments and dsRNA bands of the same size were observed.

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Example 13 - Selection and Specificity of cDNA Clones

A total of 182 clones were selected after plaque hybridization. Eighty clones with strong hybridization signals were subcloned into pBluescript SK through in vivo excision. Resulting plasmids were shown to have inserts ranging from 0.3 to 3.0 kb. A total of 20 clones with inserts of ca. 0.8 kb or larger were selected. Southern analysis of these 20 clones to radio labeled first strand cDNA probes derived from the dsRNA resulted in 15 clones with strong hybridization signals. Several of these clones were used to determine the genome sequence of the dsRNA: RSP3, RSP28, RSP94, RSP140, RSP95, and TA5. Another clone (RSP149), which was 97% similar in nucleotide sequence to RSP95, was used as one of the two probes in Northern hybridization.

Northern hybridization was employed to confirm the specific relationship of clones RSP95 and RSP149 to the isolated dsRNA. These two clones gave the strongest reaction in Southern analysis described above. Initial experiments showed that RSP95 insert hybridized with the dsRNA isolated from three accessions

^{*} Probe used was insert from cDNA clone RSP149.

[†] A faint dsRNA band could be observed on the gel after electrophoresis but no hybridization signal could be seen in Northern analysis.

[‡] Although two dsRNA bands were observed in Verduzzo 233A, they were not specific to RSP, because they were either larger or smaller than the RSP-associated 8.7 kbp dsRNA and they did not hybridize to the probe in Northern analysis.

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(Colobel 257, Seyval, and Ravat 34), from which the template dsRNAs used in cDNA synthesis were isolated. As shown in Figure 2B and indicated in Table 1, use of RSP149 insert as the probe showed that this clone hybridized with the dsRNA of ca. 8.7 kb isolated from RSP infected grapevines. Furthermore, the intensity of hybridization signals corresponded to that of the dsRNA bands observed on agarose gels stained with EtBr. Colobel 257, Seyval, Ravat 34, Grande Glabre, and Serve Villard 14-287 reacted strongly; Bertille Seyve 5563, Couderc 28-112, Joffre, and Verdelet had weak hybridization signals. The result for Ill 344-1 was not conclusive. Aminia and Canandaigua did not show visible dsRNAs or hybridization in Northern analysis. Bertille Seyve 3408, which was tested in a separate experiment, did show a ca. 8.7 kb dsRNA which hybridized to the probe from RSP149. Freedom and Verduzzo 233A, which had indexed negative for RSP on St. George, were also negative in Northern blot.

15 Example 14 - Nucleotide Sequence and Genome Structure of RSPaV-1

Six cDNA clones and three RT-PCR amplified cDNA fragments (identified as RSPA, RSPB, and RSPC) were sequenced on both strands and used to obtain the complete nucleotide sequence of a viral agent, which is shown in Figure 3A. The genome of RSPaV-1 consisted of 8726 nts excluding a poly (A) tail on the 3' end. The sequence of RSPA indicated that the 5' first base of the RSPaV-1 genome appeared to be a cytosine (C). Clone TA5, which represented the 3' end of the RSPaV-1 genome, contained a stretch of adenines (A) preceded by a cytosine.

MapDraw analysis, shown at Figure 3B, indicated that the genome of RSPaV-1 had five potential ORFs on its positive strand, while no ORFs were observed on the negative strand (data not shown). ORF1 (nt 62 to 6547 of SEQ. ID. No. 1) has a nucleotide sequence corresponding to SEQ. ID. NO. 2. ORF1 believed to encode a protein or polypeptide having a molecular weight of about 244 kDa and an amino acid sequence corresponding to SEQ. ID. No. 3. According to Lutcke et al., "Selection of AUG Initiation Codons Differs in Plants and Animals," <u>Eur. Mol. Biol. J.</u>, 6:43-48 (1987), which is hereby incorporated by reference, the start codon of ORF1 was in a favorable context: GCAAUGGC, where the "GC" after the start codon is important for initiating translation in a plant system. ORF2 (nt 6578 to 7243 of

SEQ. ID. No. 1) has a nucleotide sequence corresponding to SEQ. ID. No. 4. ORF2 is believed to encode a protein or polypeptide having a molecular weight of about 24.4 kDa and an amino acid sequence corresponding to SEQ. ID. NO. 5. The first two ORFs were separated by an intergenic region of 30 nts. ORF3 (nt 7245 to 7598 5 of SEO. ID. NO. 1) has a nucleotide sequence corresponding to SEO. ID. No. 6. ORF3 is believed to encode a protein or polypeptide having a molecular weight of about 12.8 kDa and an amino acid sequence corresponding to SEQ. ID, NO. 7. ORF4 (nt 7519 to 7761 of SEQ. ID. NO. 1), which overlapped with ORF3 by 80 nts, has a nucleotide sequence corresponding to SEQ. ID. No. 8. ORF3 is believed to encode a 10 protein or polypeptide having a molecular weight of about 8.4 kDa and an amino acid sequence corresponding to SEQ. ID. No. 9. Nine nucleotides downstream of ORF4 was the start of ORF5 (nt 7771 to 8550 of SEQ. ID. No. 1), which has a nucleotide sequence corresponding to SEO, ID, No. 10. ORF5 is believed to encode a protein or polypeptide having a molecular weight of about 28 kDa and an amino acid sequence 15 corresponding to SEQ. ID. No. 11. Downstream of ORF5 was the 3' end LJTR of 176 nts. Although computer assisted analysis indicated that two shorter ORFs may exist as alternatives to ORF1 and ORF5, neither of them were in good contexts for translation initiation.

20 <u>Example 15</u> - Comparison of the RSPaV-1 Genome with ASPV and PVM Carlavirus Genomes

The arrangement of the ORFs and the amino acid sequences of RSPaV-1 showed similarities to those of PVX (Skryabin et al., "The Nucleotide

25 Sequence of Potato Virus X RNA," Nucleic Acids Res. 16: 10929-30 (1988), which is hereby incorporated by reference), PVM (Zavriev et al., "Complete Nucleotide Sequence of Genomic RNA of the Potato M-Virus," Molecular Biology (Mosk.)

25:761-69 (1991), which is hereby incorporated by reference), and ASPV (Jelkmann, "Nucleotide Sequences of Apple Stem Pitting Virus and of the Coat Protein of a

30 Similar Virus from Pear Associated with Vein Yellows Disease and Their Relationship with Potex- and Carlaviruses," J. General Virology 75:1535-42 (1994), which is hereby incorporated by reference), with the latter two being the most similar to RSPaV-1. A representation of the sequence comparison is shown in Figure 3B and the percent identities in amino acid sequences of the ORF of RSPaV-1 and the

corresponding ORF of ASPV, PVM, and PVX are shown in Table 2 below. These analyses suggest that the ORFs of RSPaV-1 are compared with those of PVM and ASPV.

Table 2

			Triple Ge	Coat Protein				
	Region I	ORF1 Region II aa 1354-2161	Total	ORF2	ORF3	ORF4	Total	ORF5 aa142-245
ASPV	49.2	57.5	39.6	38.0	39.3	27.1	31.3	49.5
PVM	47.2	53.2	37.6	34.8	31.2	19.0	21.2	33.3
PVX	18.9	20.4	15.7	23.5	31.3	22.9	27.4	42.9

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When the total amino acid sequence of RSPaV-1 ORF1 was used for comparison, it showed 39.6% and 37.6% identities with the replicases of ASPV and PVM respectively (Table 2). These homologies were mainly found in regions I (aa 1 to 372) and II (aa 1354-2161), which are at the N and C terminal portions of the putative replicase, respectively, shown at Figures 4A and 4B. Within region I, the identities of RSPaV-1 with ASPV and PVM were 49.2% and 47.2%, respectively (Table 2). The methyltransferase domain, which is conserved in Sindbis-like superfamily of plant viruses (Rozanov et al., "Conservation of the Putative Methyltransferase Domain: A Hallmark of the "Sindbis-like" Supergroup of Positive-Strand RNA Viruses," J. General Virology 73:2129-34 (1992), which is hereby incorporated by reference), was found in this region (Figure 4A). Region II, on the other hand, showed even higher identities: 57.5% with ASPV and 53.2% with PVM (Table 2). A NTP binding motif "GXXXXGKS/T" (aa 1356 to 1363) ("X" stands for any amino acid residue), which is conserved in helicase proteins and helicase domains of eukaryotic positive strand RNA viruses (Gorbalenya et al., "A Novel Superfamily of Nucleotide Triphosphate-Binding Motif Containing Proteins which are Probably Involved in Duplex Unwinding in DNA and RNA Replication and Recombination," FEBS Letters, 235:16-24 (1988), which is hereby incorporated by reference), was found in the beginning of region II (Figure 4B). The amino acid sequences of this motif in ASPV and PVM were identical to that of RSPaV-1 except for one position. Furthermore, amino acid sequence surrounding the GDD motif, which is conserved in all RNA dependent RNA polymerases of positive strand RNA viruses (Koonin, "The

Phylogeny of RNA-Dependent RNA Polymerases of Positive-Strand RNA Viruses," J. Gen. Virology 72:2197-2206 (1991), which is hereby incorporated by reference), was located near the C terminus of the RSPaV-1 replicase protein and showed high identities to those of ASPV and PVM (Figure 4B). Other conserved residues of positive strand RNA viruses as described by Koonin, "The Phylogeny of RNA-Dependent RNA Polymerases of Positive-Strand RNA Viruses," J. Gen. Virology 72:2197-2206 (1991), which is hereby incorporated by reference, were also found in this region. Based on these information, it was concluded that ORF1 of RSPaV-1 codes for the putative replicase protein.

10 The triple gene block is a common feature of several groups of plant viruses including carlaviruses, potexviruses, and ASPV. Comparison of RSPaV-1 ORF2 with those of PVM and ASPV showed evenly distributed homologies in amino acid sequence: 38.0% identity to ASPV and 34.8% to PVM (Table 2). The N terminal region of the 24.4K protein (ORF2) contained the consensus sequence "GXGKS S/T" (aa 31 to 36) (Figure 5A), which is observed in its counterparts in carlaviruses 15 (Zavriev et al., "Complete Nucleotide Sequence of Genomic RNA of the Potato M-Virus," Molecular Biology (Mosk.) 25:761-69 (1991), which is hereby incorporated by reference) and a number of ATP and GTP binding proteins (Zimmem, "Evolution of RNA Viruses," in RNA Genetics, Holland et al., eds., CRC Press, Boca Raton, Florida, USA (1987), which is hereby incorporated by reference). The 12.8K protein 20 of RSPaV-1 encoded by ORF3 had 39.3% and 31.2% identities with its counterparts in ASPV and PVM respectively (Table 2). However, most of the matching occurred in a region from aa 29 to 62, among which 18 aa were fully conserved in all three viruses (Figure 5B). These 12-13K proteins may function in membrane binding (Morozov et al., "Nucleotide Sequence of the Open Reading Frames Adjacent to the 25 Coat Protein in Potato Virus X Genome," FEBS Letters 213:438-42 (1987), which is hereby incorporated by reference). The 8.4K protein encoded by RSPaV-1 ORF4, in contrast, showed much lower identities: 27.1% with that of ASPV and 19.0% with that of PVM (Table 2). However, four residues "TGES" (aa 38 to 41) were conserved in all three viruses (Figure 5C). In vitro studies indicated that the analogous 7K 30 protein of PVM may bind to single or double stranded nucleic acids (Gramstat et al., "The 12 kDa Protein of Potato Virus M Displays Properties of a Nucleic Acid-Binding Regulatory Protein," FEBS Letters, 276:34-38 (1990), which is hereby

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A sequence similarity search in a DNA database revealed identities

incorporated by reference) and to plasma membrane (Morozov et al., "In vitro Membrane Binding of the Translation Products of the Carlavirus 7-kDa Protein Genes." Virology 183:782-85 (1991), which is hereby incorporated by reference).

between the putative protein encoded for by RSPaV-1 ORF5 to the coat proteins 5 (CPs) of several groups of plant viruses, indicating that RSPaV-1 ORF5 may code for the coat protein. MegAlign analysis revealed that RSPaV-1 ORF5 had 31.3% and 21.2% identities with the CPs of ASPV and PVM, respectively (Table 2). Most of the identities were found in the C terminal portion of the coat proteins (aa 142 to 245 for RSPaV-1), while the N terminal portions were quite variable in the numbers and 10 sequences of amino acid residues. When the C terminal portion of RSPaV-1 CP was compared to the corresponding regions of ASPV and PVM, it showed 49.5% and 33.3% identities with ASPV and PVM, respectively (Table 2). In addition, the "RR/QX-XFDF" motif was found in the central region of RSPaV-1 CP (Figure 5D). This motif is conserved in the CPs of positive strand RNA viruses with filamentous 15 morphology and were reported to be involved in salt bridge formation (Dolja et al., "Phylogeny of Capsid Proteins of Rod-Shaped and Filamentous RNA Plant Virus: Two Families with Distinct Patterns of Sequence and Probably Structure Conservation," Virology, 184:79-86 (1991), which is hereby incorporated by reference). Therefore, it is believed that ORF5 encodes a putative coat protein. 20

MegAlign analysis, shown in Figures 6A and 6B, revealed that the 3' UTR of RSPaV-1 is more similar to that of PVM than to that of ASPV. For example, in a 75 nts stretch, RSPaV-1 had 68% identity with PVM. Within this region, 21 consecutive nucleotides were identical between these two viruses. The significance of this conservation in nucleotide sequence remains to be explored. In contrast, the 5' UTR of RSPaV-1 did not reveal significant similarities with those of PVM and ASPV.

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It has been have shown that an 8.7 kbp dsRNA is consistently associated with grapevines that indexed positively on St. George for RSP. Sequence analyses of the dsRNA provide evidence that a virus is involved in RSP, which has now been named RSPaV-1. The complete nucleotide sequence of RSPaV-1 was determined from overlapping cDNA clones and RT-PCR-amplified cDNA fragments generated from the dsRNA. The RSPaV-1 genome has five ORFs coding for the

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putative replicase (ORF1), the triple gene block (ORF2-4), and the CP (ORF5). The existence of these ORFs and their potential to code for structural and non-structural viral proteins were further supported by the identification of conserved motifs which are the signatures of various viral proteins.

This work confirms and extends the findings of Walter and Cameron ("Double-stranded RNA Isolated from Grapevines Affected by *Rupestris* Stem Pitting Disease," Am. J. Enology and Viticulture 42:175-79 (1991), which is hereby incorporated by reference), and Azzam and Gonsalves ("Detection of dsRNA in Grapevines Showing Symptoms of Rupestris Stem Pitting Disease and the Variabilities Encountered," Plant Disease 75:960-64 (1991), which is hereby incorporated by reference), who observed a major dsRNA species of about 8.0-8.3 kbp in RSP-infected grapevines. In addition, such work also observed a smaller dsRNA of ca. 6.6 kbp. A dsRNA of similar size was also observed here, but it was consistently detected in only Colobel 257 and Seyval. The relationship, if any, of this smaller dsRNA to RSP remains to be determined. The small dsRNA of ca. 0.359 kbp, which Monette et al. ("Double-stranded RNA from Rupestris Stem Pitting-Affected Grapevines," Vitis 28:137-44 (1989), which is hereby incorporated by reference) isolated from RSP-infected grapevines growing in tissue culture, was not observed.

Electron microscopy evidence also suggests that RSP is caused by filamentous virus(es). Tzeng et al. ("Anatomical and Tissue Culture Studies of Rupestris Stem Pitting-Affected Grapevines," Botan. Bulletin of Acad. Sinica (Taipei) 34:73-82 (1993), which is hereby incorporated by reference) observed flexuous filamentous virus aggregates in the phloem parenchyma cells of young shoots of Sylvner grapevines that had indexed positively for RSP. Monette and Godkin ("Detection of Capillovirus-like Particles in a Grapevine Affected with Rugose Wood," Vitis 34:241-42 (1995), which is hereby incorporated by reference) observed a filamentous virus in Sauvignon blanc infected by RSP and LNSG. The relationship of these virus particles to RSP disease remains to be studied.

Evidence suggests that the cDNA library generated from the isolated dsRNA templates is not homogeneous for only RSPaV-1. During the process of

sequencing cDNA clones, several clones (e.g., RSP47-4 and RSP158) were identified with high, but not identical, sequence similarities to RSPaV-1.

RSPaV-1 has the most similarities to ASPV, which has not yet been grouped into a virus genus. Both viruses have the same genome organization and their ORFs code for putative proteins of similar sizes, except that the coat protein of 5 ASPV is significantly larger (44 kDa) than that of RSPaV-1 (28 kDa). Comparisons of RSPaV-1 with PVM carlavirus show some similarities in genome organization except that RSPaV-1 lacks ORF6 which is located at the 3' end of PVM genome. Although the genome organization of RSPaV-1 is similar to PVX potexvirus, the latter has a much smaller putative replicase. RSPaV-1 has no relation to grape viruses 10 whose genomes have been sequenced so far. The closest possibilities, GVA (Minafra et al., "Grapevine virus A: Nucleotide Sequence, Genome Organization, and Relationship in the Trichovirus Genus," Arch. Virology 142:417-23 (1997), which is hereby incorporated by reference) and GVB (Saldarelli et al., "The Nucleotide Sequence and Genomic Organization of Grapevine Virus B," J. General Virology 15 77:2645-52 (1996), which is hereby incorporated by reference), have different genome structures than RSPaV-1.

Example 16 - Specific and Universal Primers and the Detection of Different Strains of RSPaV by Reverse Transcription-Polymerase Chain Reaction (RT-PCR)

Among the 138 grapevine entries collected, 25 indexed negatively and 93 indexed positively for RSP on St. George, while the others were not indexed (see Tables 3-7 below). Symptoms induced by RSP on the woody cylinder of St. George after graft inoculation with chip-buds can be divided into two types. The first type is called "specific", that is, pits and/or grooves being restricted to the area on the woody cylinder below the inoculation sites. The other is called "nonspecific", that is, pits and/or grooves being present above, around, and below the inoculation sites.

Table 3

	I abic 5			
Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Almeria K3 P 661	1483-13D1	•	•	С
Auxerrois CL 56	658-1A2	-	-a	С
Auxerrois CL 56	658-1A1-1A2	-	-	С
GM 32458	604-8A2-2A2	-	-	С
GM 7117-10	1347-16A1	-	-a	С

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Table 3

Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Italia	1186-5B1	. •	-	С
Pslanka (H)	23-10A2-2A2	-	-	·C
Ventura (V. 51061) (H)	1166-2A1	-	-	С
Verdelet (H)	1170-3C2-2S1	-	-	С
Verduzzo (V)	233A	-	-	I
Vivant (V. 63331) (H)	1166-3A1	-	· -	С
Control				
Sauvignon Blanc (V)	AV-4 #2	-	-a	Ū

Symbols:

Table 4

Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Aragonez (Temperanillo)	238	•	+	P
Albalonga	1058-4A2-2A1	-	+	С
Cabernet Franc (V)	147A	-	+	I
Chardonnay (V)	80A	•	+	I
Ehrenfelser PM 1 (V)	1169-1A1	-	+	C
Freedom (H)	PI 588370	-	+a	U
Harslevellu P 679	1483-2B1		+	С
Heroldrebe	1318-2A1	•	+	С
Malvasia Fina	340	. -	+	P
Perle of Zala	1407-5A1	-	. +	С
Refosco (V)	181A	-	. +	I
San Giovese Brunello CL BBS 11	1497-2A1	-	+	С
Touriga Francesa	313	•	+ -	P

Symbols:

Table 5

ID	Index St.G	RT-PCR	Source
1058-4A2-1A2	+	+	С
PI 588306	+	+	U
CL 245	+	+	P
350	+	+	P
658-1A1	+	+	С
1407-1A1	+	+	С
GVIT 348	+	+b	U
PI 181647	+	+a	U
Q1378-1	+	+ b	С
1240-1A1	+	+a	С
1586-17P3	+	+	С
1078-1A1	+	+ -	С
151A	+	+	I
124A	+	+	I
Q390-13	+	+b	С
Q661-4	+	+b	С
	1058-4A2-1A2 PI 588306 CL 245 350 658-1A1 1407-1A1 GVIT 348 PI 181647 Q1378-1 1240-1A1 1586-17P3 1078-1A1 151A 124A Q390-13	1058-4A2-1A2 + PI 588306 + CL 245 + 350 + 658-1A1 + 1407-1A1 + GVIT 348 + PI 181647 + Q1378-1 + 1240-1A1 + 1586-17P3 + 1078-1A1 + 151A + 124A + Q390-13 +	1058-4A2-1A2

V., Vitis vinefera; R., Vitis riparia; H., hybrid; C., Canada; I., Italy; U., USA; P., Portugal; a, tested by RSP149F1/R1 and 95F1/R1 and results agree to each other; b, tested by 95F1/R1 only

V., Vitis vinefera; R., Vitis riparia; H., hybrid; C., Canada; I., Italy; U., USA; P., Portugal; a, tested by RSP149F1/R1 and 95F1/R1 and results agree to each other; b, tested by 95F1/R1 only

Table 5

Cultivar/Accession		1 able			
Chardonnay (V)	Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Chardonnay (V)	Chardonnay CL 116 (V)	1021-13A2	+		С
Chardonnay (V) 72A + +b I Chardonnay (V) 73A + +b I Chardonnay (V) 83A + +b I Chardonnay (V) 945-81 + +b C Cha		128B	+	+b	I
Chardonnay (V)	Chardonnay (V)	72A	+	+b	I
Chardonnay (V) 83A + + 1 Chazan CL 538 1346-6A1 + + 1 Chazan CL 200 1555-6A1 + + 1 Colobel 257 (Seibel 8357) (H) P1 588062 + + 1 De Chaunac S9549 (H) Q659-1 + 1 De Chaunac S9549 (H) Q659-1 + + 1 De Camay Precoce 1 500-2A1 + + 1 De Camay Precoce 1		73A	+	+b	I
Charian Blanc CL 220 1555-6A1 + +a C Choinbel Blanc CL 220 1555-6A1 + +a U Couder 28-112 (H) PI 588062 + +a U Durella 3 1586-13Pl + +b C Charlanac 89549 (H) Q659-1 + +b C Durella 3 1586-13Pl + +C Camay Precoce Egri Csillagok-30 1407-3A1 + +C Camay Precoce 1500-2A1 + +c C Gamay Precoce 1500-2A1 + +c C GM 31875 782-18A1 + +c C GM 31875 782-18A1 + +c C GM 32458 782-21B1 + +C C GM 497-4 1347-7A1 + +C C GM 497-4 1347-7A1 + +C C GM 7116-10 1362-4A1 + +C C GM 7116-10 1362-6A1 + +C C GM 7116-10 1362-6A1 + +C C GM 7116-10 GM 7117-13 147-17A2 + +C C GM 7116-10 GM 7117-13 147-17A2 + +C C GM 7117-13 147-17A2 +C C GM 7117-13 C GM 7117-13 C GM 7117-13 C GM 7117-13 C GM 7117-		83A	+	+	I
Colobel 257 (Seibel 3357) (H) PI 588062 + +a U Couder 28-112 (H) PI 588248 + +a U De Chaunac 89549 (H) Q659-1 + +b C Durella 3 1586-1391 + +b C Esgana cao 276 + + + P P Esgana cao 276 + + + P P Esgana cao 276 + + + C Gamay Precoce 1500-2A1 + + + C C Gamay Precoce 1500-2A1 + + + C C GM 31875 782-18A1 + + a C GM 32458 604-8A1 + + C C GM 31875 782-18A1 + + C C GM 32458 782-21B1 + + C C GM 4047-7 1347-7A1 + + C C GM 6497-4 1347-7A1 + + C C GM 7116-10 1362-4A1 + + C C GM 7116-10 1362-4A1 + + C C GM 7117-13 1347-17A2 + + C C GM 7116-10 1362-4A1 + + C C GM 7117-13 1347-17A2 + + C C GM 7117-13 1347-17A2 + + C C GM 7117-13 1347-17A2 + + C C GM 7117-13 1347-17A1 + + C C GM 7117-14 +		1346-6A1	+	+a	. C
Coudere 28-112 (H) PI 588248 + +a U De Chamnac S9549 (H) Q659-1 + +b C Durella 3 1586-13P1 + + + C Esgana cao 276 + + + P Esgana cao 276 + + + P Esgri Csillagok-30 1407-3A1 + + C Gamay Precoce 1500-2A1 + + C GM 31875 782-18A1 + +a C GM 31875 782-18A1 + +a C GM 32458 604-8A1 + + C GM 32458 782-21B1 + + C GM 6497-4 1347-14A1 + + C GM 7116-10 1362-4A1 + + C GM 7116-10 1362-4A1 + + C GM 7117-13 1347-17A2 + + C GM 7117-13 1347-17A2 + + C Grande Glabre (R) 279897 + +a U Gyongyriziling 1407-4A1 + + C ILL 344-1 (H) GVIT 588 + +a U Joffre (Xuhlmann 187-1) (H) GVIT 588 + +a U Joffre (Xuhlmann 187-1) (H) GVIT 381 + +a U Malvasia (V) 153A + 1 I Malvasia (V) 153A + 1 I Malvasia (V) 153A + 1 I Malvasia (V) 1236-17A1 + C Moureto 87 + P Moureto 87 + P Moureto 187 + P Moureto 187 + P Pinot Chardonny CL 270 (V) 949-3A2 + + C Pinot Chardonny CL 277 (V) 949-8B1 + + C Pinot Grigio (V) 108A + + I Pollux B6-18 1357-4A1 + + C San Giovese Brunello CL E BS 4 1497-3B1 + + C Serval Blanc (L 98 (V) 1555-2A1 + + C Serval Blanc (L 98 (V) 158209 + + a C Serval Blanc (L 98 (V) 158209 + + a C Serval Blanc (L 98 (V) 158209 + + a C Serval Blanc (Seyve Villard 5-276) (H) Seyve Villard 3160 (H) PI 181630 + + a U Verdelto (V) 4044 + + b C Verdelto (PI 186260 + + a U Verduzzo (V) 222A + + b I Verduzzo (V) 222A + + b I	Chenin Blanc CL 220	1555-6A1	+	+	C
Couder 28-112 (H) PI 588248 + +a U De Channac 99549 (H) Q659-1 + +b C Durella 3 1586-13P1 + +b C Eagana cao 276 + + P Esgana cao 276 + + P Egri Csillagok-30 1407-3A1 + + C Gamay Precoce 1500-2A1 + +c C GM 31875 782-18A1 + +a C GM 32458 604-8A1 + +C C GM 32458 782-21B1 + +C C GM 6497-4 1347-14A1 + +C C GM 7116-10 1362-4A1 + +C C GM 7116-10 1362-4A1 + +C C GM 7117-13 1347-17A2 + +C C GM 7116-10 Grande Glabre (R) 279897 + +a U Gyongyrizling 1407-4A1 + +C C GYongyrizling 1407-4A1 + +C C GLAGH (Kuhmann 187-1) (H) GVIT 658 + +a U Joffre (Kuhmann 187-1) (H) GVIT 381 + +b C Malvasia (V) 153A + U Malvasia (V) 153A + I Malvasia (V) 153A + I Malvasia (V) 1236-17A1 + +C C Moureto 87 + + P Moureto 187 + + C Pinot Chardonnay CL 76 (V) 949-3A2 + +a C Pinot Chardonny CL 277 (V) 949-8B1 + +C C Pinot Chardonny CL 277 (V) 949-8B1 + +C C Pinot Grigio (V) 108A + +b I Pollux B6-18 1357-4A1 + + C San Giovese Brunello CL E BS 4 1497-3B1 + +a C Semillon CL 299 (V) 1555-2A1 + +a C Semillon CL 299 (V) 1555-7A1 + +a C Semillon CL 299 (V) 1555-7A1 + +a C Seywe Villard 3-60 (H) PI 181630 + +a U Verdelto (Preduzzo (V) 22A + +b I Verduzzo (V) 22A + +b I Verduzzo (V) 222A + +b I		PI 588062	+	+a	U
De Chaunac S9549 (H) Q659-1 + +b C Durella 3 1586-13P1 + + C Esgana cao 276 + + P Esgana cao 1407-3A1 + + C Gamay Precoce 1500-2A1 + + C GM 31875 782-18A1 + + C GM 32458 604-8A1 + + C GM 32458 782-21B1 + + C GM 6417-7 1347-7A1 + + C GM 6417-7 1347-7A1 + + C GM 7116-10 1362-4A1 + + C GM 7116-10 1362-4A1 + + C GM 7116-10 1362-4A1 + + C GM 7116-10 1367-4A1 + + C GM 7116-10 1367-4A1 + + C Grande Glabre (R) 279897 + + a U Gyongyriziling 1407-4A1 + + C Grande Glabre (R) GVIT 658 + a U Joffre (Kuhlmann 187-1) (H) GVIT 381 + + a U Koret (H) GVIT 381 + + a U Koret (H) GVIT 381 + + C Malvasia (V) 153A + I Malvasia (V) 161A + + I Merlot CL 447 (V) 1236-17A1 + + C Moureto 87 + + P Moureto 87 + + P Moureto 96 + + P Muscat De Hambourg CL 202 1346-5A1 + + C Perie of Csaba Q806-1 + + b C Prinot Grigio (V) 108A + + I Pinot Grigio (V) 108A + + I Pinot Grigio (V) 108A + + B Pinot Grigio (V) 108A + + B Pinot Grigio (V) 108A + + C Pinot Grigio (V) 108A + + C Ravat 34 Pi588247 + + C Pinot Grigio (V) 108A + + C Ravat 34 Pi588247 + + C Ravat 34 Pi588247 + + C Refosco (V) 190A + + C Refosco (V) 195A + + C Refos		PI 588248	+	+a	ប
Durella 3		Q659-1	+	+b	С
Esgna cao 276 + + + P Egri Csillagok-30 1407-3A1 + + + C Gamay Precoce 1500-2A1 + + + C GM 31875 782-18A1 + + + A C GM 31875 782-18A1 + + + C GM 32458 604-8A1 + + C GM 32458 782-21B1 + + C GM 6417-7 1347-7A1 + + C GM 6417-7 1347-7A1 + + C GM 6417-7 1347-7A1 + + C GM 7116-10 1362-4A1 + + C GM 7116-10 1362-4A1 + + C GM 7117-13 1347-17A2 + + C Grande Glabre (R) 279897 + + A Gyongyriziling 1407-4A1 + + C Grande Glabre (R) GVIT 658 + A U Joffre (Kuhimann 187-1) (H) GVIT 381 + + A U Joffre (Kuhimann 187-1) (H) GVIT 381 + + A U Koret (H) Q1179-7 + + b C Malvasia (V) 161A + + 1 Merlot CL 447 (V) 1236-17A1 + + C Moureto 87 + + P Moureto 96 + + P Muscat De Hambourg CL 202 1346-5A1 + + C Prinot Chardonny CL 76 (V) 949-3A2 + + A Pinot Chardonny CL 77 (V) 949-8B1 + C Prinot Chardonny CL 77 (V) 949-8B1 + + C Pinot Grigio (V) 104A + + b I Pinot Grigio (V) 108A + + b I Pinot Grigio (V) 108A + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588247 + + A PSlanka (H) 23-10A2 + + C Ravat 34 PI 588246 + A PSlanka (H) + A PSlanka			+	+	
Egri Csillagok-30		276	+	+	P
Gamay Precoce 1500-2A1			+	+	
GM 3 1875			+	+	
GM 32458 604-8A1 + + C GM 32458 782-21B1 + + + C GM 6497-4 1347-7A1 + + + C GM 6497-4 1347-7A1 + + + C GM 7116-10 1362-4A1 + + + C GM 7117-13 1347-17A2 + + + C GM 7116-10 1362-4A1 + + + C GM 7117-13 147-17A2 + + + C ILL 344-1 (H) GVIT 588 + + a U Joffre (Kuhlmann 187-1) (H) GVIT 381 + + a U Koret (H) Q1179-7 + + +b C Malvasia (V) 153A + + 1 Malvasia (V) 153A + + 1 Malvasia (V) 153A + + 1 Malvasia (V) 161A + + C Moureto 87 + + P Moureto 87 + + P Moureto 87 + + P Muscat De Hambourg CL 202 1346-5A1 + + C Perle of Csaba Q806-1 + + b C Perle of Csaba Q806-1 + + b C Prinot Chardonnay CL 76 (V) 949-3A2 + + a C Pinot Chardonnay CL 77 (V) 949-8B1 + + C Pinot Grigio (V) 104A + + b I Pinot Grigio (V) 114A + + 1 Pollux B6-18 1357-4A1 + + C Ravat 34 P158247 + + a C San Giovese Brunello CL E BS 4 1497-3B1 + + C San Giovese Brunello CL E BS 4 1497-3B1 + + C Semillon CL 299 (V) 1555-2A1 + + a C Semillon CL 299 (V) 1555-2A1 + + a C Semillon CL 299 (V) 1555-7A1 + + a C Semillon CL 299 (V) 1555-7A1 + + a C Semillon CL 297 (H) P1 181630 + + a U Verdelto V Verdelho 274 + + b I Verduzzo (V) 222A + + b I Verduzzo (V) 222A + + b I			+	+a	C
GM 32458 782-21B1 + + C GM 6497-7 1347-7A1 + + + C GM 6497-4 1347-14A1 + + + C GM 6497-4 1347-14A1 + + + C GM 7116-10 1362-4A1 + + + C GM 7117-13 1347-17A2 + + + C Grande Glabre (R) 279897 + +a U Gyongyriziling 1407-4A1 + + + C ILL 344-1 (H) GVIT 568 + +a U Joffre (Kuhlmann 187-1) (H) GVIT 381 + +a U Koret (H) Q1179-7 + + +b C Malvasia (V) 161A + + I Malvasia (V) 161A + + I Merlot CL 447 (V) 1236-17A1 + + C Moureto 87 + + P Moureto 96 + + P Muscat De Hambourg CL 202 1346-5A1 + + C Perle of Csaba Q806-1 + +b C Pinot Chardonnay CL 76 (V) 949-3A2 + +a C Pinot Chardonnay CL 277 (V) 949-8B1 + + C Pinot Grigio (V) 104A + +b I Pinot Grigio (V) 108A + +b I Pinot Grigio (V) 108A + +b I Pinot Grigio (V) 114A + + C Ravat 34 P 1588247 + +a U Refosco (V) 190A + +? I Refosco (V) 190A + +2 C San Giovese Brunello CL E BS 4 1497-3B1 + + C Semillon CL 299 (V) 1555-7A1 + +a C Sewillon CL 299 (V) 1555-7A1 + +a C Sewillon CL 299 (V) 1558-309 + +a U Seyve Villard 3160 (H) P1181630 + +a C Sevyal Blanc (Seyve Villard 3160 (H) P1181630 + +a U Verdelbo 274 + +b I Verduzzo (V) 222A + +b I Verduzzo (V) 222A + +b I			+	+	Ċ
GM 6417-7			+	+	Č
GM 6497-4					Ċ
GM 7117-13					Č
GM 7117-13					Č
Grande Glabre (R) 279897 + + +a U Gyongyriziling 1407-4A1 + + C ILL 344-1 (H) GVIT 658 + + +a U Joffre (Kuhlmann 187-1) (H) GVIT 381 + +a U Koret (H) Q1179-7 + +b C Malvasia (V) 153A + + I Malvasia (V) 161A + + 1 Merlot CL 447 (V) 1236-17A1 + + C Moureto 87 + + P Moureto 96 + + P Muscat De Hambourg CL 202 1346-5A1 + + C Perle of Csaba Q806-1 + +b C Pinot Chardonnay CL 76 (V) 949-3A2 + +a C Pinot Chardonny CL 277 (V) 949-8B1 + + C Pinot Grigio (V) 104A + +b I Pinot Grigio (V) 108A + +b I Pinot Grigio (V) 114A + + I Pollux B6-18 1357-4A1 + + C Ravat 34 PI 588247 + +a U Refosco (V) 190A + +7 Refosco (V) 195A + +1 Refosco (V) 195A + +1 Refosco (V) 195A + + C San Giovese Brunello CL E BS 4 1497-3B1 + + C Semillon CL 299 (V) 1555-7A1 + +a C Semillon CL 299 (V) 1558246 + +a U Seyve Villard 5-276 (H) Seyve Villard 14-287 (H) PI 181630 + +a U Verdelbo 274 + +b I Verduzzo (V) 222A + +b I					Č
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Pollux B6-18	Pinot Grigio (V)	114A	+	+	
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verauzzo (V) 239A + + 1					
	verauzzo (V)	239A	+	+	1

Table 5

Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Vidal Blanc	1200-5A1	Index of o		Source
Weiser Burgunder	Q782-40	+	+a +b	C
3309 C	330-4A1	+	+	C
420 A	1483-4A1	+	+	Č
7542	Q1386-1	+	+ b	Č
Pinot Noir (V)	1186-9A2	+	+a	Č
Thompson Seedless (V)	RSP105	+	+a	Ū

Symbols:

Table 6

	lable 6	· · · · · · · · · · · · · · · · · · ·		
Cultivar/Accession	ID	Index St.G	RT-PCR	Source
Aligote	Q637-2B2	+	-b	С
Aragonez (Temperanillo)	232	+	•	P
Canandaigua (H)	GVIT 566	+	-a	ប
Challenger (H)	Q1338-1	+	-b	Ċ
Fercal CL 242	1551-4A1	· +	-a	Č
GM 7746-6	1362-6A1	+	-	Ċ
Gravesac CL 264	1551-3A1	+	-a	Ċ
Honey Red	1339-6A1	+	.=	Č
Kee-Wah-Din (H)	1278-1A1	+	•	Ċ
Periquita	72	+	-	P
Tajoznyt Izumrud (H)	Q2-2	+	-b	CCCCPCC
Thurling	1047-4A2-1A2	+	•	č
Verdelet	1170-3D2-2A1	+ .	-	Ċ.
5BB CL 114	1236-2A1	. +	-	Č
Alphonse Lavalle		NI	+	· T
Ancellotta		NI	+	Ī
Chardonnay (V)	127	NI	+	Î
Kober 5BB?	100	NI	+	ī
Moscato d'Adda	7	NI	+	Î
Periquita	624	NI	+	P
Periquita	633	NI	+	P
Riesling (V)	3	NI	+	Ī
Seyval (H)	Peterson	NI	+	Ū
Теттапо	1/1/3/K	NI	+	Ĭ
Thurling	1047-4A2-2A1	NI	•	ċ
Tocai Rosso 19	1586-21P4	NI	• +	č
Trebbiano Toscano	67	NI	-	Ĭ
Vidal	Peterson	NI	+	Ū

Symbols:

V., Vitis vinefera; R., Vitis riparia; H., hybrid; C., Canada; I., Italy; U., USA; P., Portugal; a, tested by RSP149F1/R1 and 95F1/R1 and results agree to each other; b, tested by 95F1/R1 only.

V., Vitis vinefera; R., Vitis riparia; H., hybrid; NI, not indexed; C., Canada; I., Italy; U., USA; P., Portugal;

a, tested by RSP149F1/R1 and 95F1/R1 and results agree to each other; b, tested by 95F1/R1 only

Table 7

Cultivar/Accession	ID	Index St.G	RT-PCR	Source
V. acerifolia	PI 588448	NI	-	U
V. acerifolia	PI 588449	NI	-	U
V. cinerea	PI 588446	NI	•	U
V. monticola	PI 588454	NI	-	U
V. riparia	PI 495622	NI	-	U
V. sp. yenshanesis	PI 588421	NI	•	U

Symbols:

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V., Vitis vinefera; R., Vitis riparia; H., hybrid; NI, not indexed; C., Canada; I., Italy; U., USA; P., Portugal;

a, tested by RSP149F1/R1 and 95F1/R1 and results agree to each other; b, tested by 95F1/R1 only

Among the 93 RSP-infected grapevines, 79 (85%) produced cDNA fragments of expected sizes in repeated RT-PCR using RSP149F1/R1 primers (SEQ. ID. Nos. 49 and 50) and/or RSP95F1/R1 primers (SEQ. ID. Nos. 47 and 48), while the other 14 were negative (see Tables 5 and 6). Interestingly, 12 of 14 (85.7%) grapevine accessions which were not indexed for RSP also produced cDNA fragments of expected size in RT-PCR (see Table 6). Sauvignon blanc (healthy control) was negative in repeated RT-PCR (see Table 3).

Results of RT-PCR for grapevines indexed negatively for RSP were surprising (see Tables 3 and 4). While 11 were negative in RT-PCR tests (excluding Sauvignon blanc healthy control), the other 13 produced cDNA fragments of expected sizes.

Since RSPaV-1 was detected not only from grapevines which indexed positively for RSP but also from some of the grapevines indexed negatively for RSP, a search for more healthy materials for RT-PCR tests became necessary. As the majority of plant viruses do not pass on through seeds, grapevine seedlings are probably free of RSPaV-1. Based on this assumption, six seedlings from five *Vitis* species were included in RT-PCR (see Table 7). None of them produce cDNA of expected size in RT-PCR using RSP149R1/F1 primers (SEQ. ID. Nos. 49 and 50).

The data described above (and shown in Tables 3-7) indicate that RSPaV-1 is closely associated with RSP and that it is likely the causal agent of RSP. RT-PCR detected RSPaV-1 specific sequences from most of the RSP-infected grapevines collected from a wide range of viticultural regions of the world. Among the 93 grapevine accessions indexed positively for RSP on St. George, 85% were positive in RT-PCR (see Table 5). The data also suggests that RT-PCR has the

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potential to be used as a standard method for diagnosing RSP. This method is advantageous over the biological indexing on indicator St. George, because it is simpler, quicker, and more sensitive.

RT-PCR did not detect RSPaV-1 sequences from 14 of the grapevine accessions indexed positively for RSP (see Table 6). The discrepancy between RT-PCR and indicator indexing can be attributed to the existence in grapevines of different viruses or strains of the same virus which may all induce similar pitting and/or grooving symptoms on St. George upon graft-inoculation. It is believed these agents are only slightly different from RSPaV-1 at the level of their nucleotide sequences, but significant enough to hinder them from being detected by RT-PCR using RSPaV-1 specific primers.

It is likely that many RSPaV strains have genomes with nucleotide sequences that are highly similar to the nucleotide sequence of the RSPaV-1 genome. Evidence that supports this hypothesis includes the finding of a highly conserved region of ca. 600 bps among the nucleotide sequences of RSPaV-1 (type strain) and seven other cDNA clones, as shown in Figure 9. The nucleotide sequence identities of these strains to RSPaV-1 (type strain) range from 83.6% to 98.4%. If oligonucleotides are chosen which are conserved among all these strains (i.e., with one or only a few mismatches), then the oligonucleotides should function as universal primers, allowing all of the strains to be detected by RT-PCR. Based on this theory, a primer pair (BM98-3F/BM98-3R) can be designed to amplify a DNA fragment of 320 bps from all these clones. BM98-3F has a nucleotide sequence corresponding to SEQ. ID. No. 51 as follows:

25 GATGAGGTCCAGTTGTTTCC

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BM98-3R has a nucleotide sequence corresponding to SEQ. ID. No. 52 as follows:

ATCCAAAGGACCTTTTGACC

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Primers BM98-3F/BM98-3R can be used in RT-PCR to test further some of the grapevine samples which were negative for RSPaV in RT-PCR using RSP95F1/RSP95R1 primers (SEQ. ID. Nos. 47 and 48, respectively) or RSP149F1/RSP149R1 primers (SEQ. ID. Nos. 49 and 50, respectively). Results show that 6 of the 9 samples included were positive for RSPaV in RT-PCR using

BM98-3F/BM98-3R primers. This indicates that these universal primers can be used to achieve even higher detection rates.

Another pair of primers (BM98-1F/BM98-1R) can be designed in a way that they can amplify DNA of 760 bps from RSPaV-1, RSP47-4, and RSP158.

5 BM98-1F has a nucleotide sequence corresponding to SEQ. ID. No. 53 as follows:

CTTGATGAGTACTTGTC

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BM98-1R has a nucleotide sequence corresponding to SEQ. ID. No. 54 as follows:

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GCAAGGATTTGGATGGC

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Other "universal primers" can be designed manually or with computer programs (such as PrimerSelect) in the same way so that they contain conserved regions of nucleotide sequences for different strains of RSPaV-1.

RT-PCR detected RSPaV-1 sequences from 54% of grapevines negative for RSP as judged by indexing on St. George (see Tables 3 and 4). Several possibilities may account for this discrepancy. First, RT-PCR is much more sensitive than indicator indexing. Virus(es) of extremely low concentration may not induce visible symptoms on St. George within the standard indexing period, while they can be detected by RT-PCR. Second, judging indexing results can, in some cases, be very subjective. For example, it is very difficult to reach a conclusion on whether a grapevine is infected with RSP when only one or a few small pits are present on the woody cylinder of St. George. Third, uneven distribution of virus(es) within grapevines and the relatively limited number of replicates of St. George indicators may result in the failure to detect RSP-infection.

RSP seems to be widespread in different types of grapevines including V. vinifera, hybrids, V. riparia, and rootstocks. It occurs in a wide range of geographic regions including North America, Europe, Australia, and possibly many other countries as well. Testing grapevines from other areas of the world using RSPaV-1 specific primers will provide definitive information on the exact distribution of RSP throughout the world. It is also interesting to investigate whether RSP is transmitted by any vectors in nature.

RSP is a disease under quarantine in Washington and New York of the USA. Since this work and the work of others (Golino and Butler, "A Preliminary

Analysis of Grapevine Indexing Records at Davis, California," in Proceedings of the 10th Meeting of the ICVG, pp. 369-72, Rumbos et al., eds., Volos, Greece (1990); Azzam and Gonsalves, "Detection of dsRNA in Grapevines Showing Symptoms of Rupestris Stem Pitting Disease and the Variabilities Encountered," Plant Disease,

5 75:96-964 (1991); Garau, "Kober Stem Grooving and Grapevine Virus A: A Possible Relationship," in Extended Abstracts of the 11th Meeting of the International Council for the Study of Viruses and Virus Diseases of the Grapevine, p. 54, Montreux, Switzerland (1993); Credi, "Characterization of Grapevine Rugose Wood Sources from Italy," Plant Disease, 82:1288-92 (1997), all of which are hereby incorporated by reference) showed that RSP is so wide-spread, it is questionable whether or not RSP should be kept under plant quarantine any longer. The devlopment and advance of rapid diagnostic methods will also allow us to investigate on the economic damage caused by RSP.

According to Goheen ("Rupestris Stem Pitting," in Compendium of 15 Grape Diseases, p. 53, Pearson and Goheen, eds., American Phytopathological Society Press, St. Paul, Minnesota, USA (1988), which is hereby incorporated by reference), RSP is a disease which induces, after graft-inoculation with a chip bud from an infected grapevine, a row of small pits on the woody cylinder of St. George below the point of inoculation. This definition may not be comprehensive. Indexing record indicated that two types of stem pitting (specific vs. nonspecific) were often 20 observed on the woody cylinder of St. George upon graft inoculation with chip buds. For example, among 16 RSP-positive grapevines collected from Canada in 1995, eight developed specific type symptoms, while the others produced nonspecific symptoms. Credi ("Characterization of Grapevine Rugose Wood Sources from Italy," 25 Plant Disease, 82:1288-92 (1997), which is hereby incorporated by reference) also observed these two types of stem pitting in his indexing work. However, from the primers used in RT-PCR, as described above, RSPaV-1 was detected in grapevines showing both types of symptoms on St. George.

Thus, RT-PCR detected RSPaV-1 sequences from a wide range of
grapevines collected from a number of major grapevine growing countries. The data
clearly suggest that RSPaV-1 is closely associated with *Rupestris* stem pitting of
grapevines and that it is likely the causal virus of RSP. Use of "universal" primers
which can detect multiple agents which are highly similar to RSPaV-1 in nucleotide

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adjuvant.

sequences would improve the detection rate by RT-PCR. In addition, antibodies produced against bacteria-expressed coat proteins of RSPaV-1 will help in finding the viral particles from RSP infected grapevines and in rapid detection of RSP.

5 Example 17 - Southern Hybridization

To confirm the specificity of the RT-PCR products to RSPaV-1,

Southern blot hybridization was conducted using 32P labeled probe specific to

RSPaV-1. As shown in Figure 7, the Southern blot hybridization confirmed the results

of the RT-PCR in each of the tested samples. Specifically, cDNA fragments amplified
by RT-PCR from 16 selected RT-PCR positive samples hybridized with the probe.

Example 18 - Constructing Expression Systems, Expression of a Fusion Protein Containing the RSPaV-1 Coat Protein, Production of Antibodies Against the Fusion Protein and Their Use in Detecting RSPaV-1 from Grapevines

The coat protein gene (SEQ. ID. No. 10) of RSPaV-1 was cloned into the EcoRI and HindIII sites of the polylinker region of a protein expression vector pMAL-c2 which, upon induction by inducer IPTG, produces a fusion protein containing maltose binding protein (MBP) and the coat protein of RSPaV-1. The fusion protein of expected size (ca. 71 KDa) was produced in *E. coli* bacteria after induction with IPTG. This fusion protein was purified through affinity chromatography using an amylose column. Purified fusion protein was used as an antigen to immunize a rabbit (by subcutaneous injection along the back) with the following scheme:

first injection, 400 μ g fusion protein in 0.5 ml column buffer with Freund's complete adjuvant; second injection, 100 μ g of protein in 0.5 ml column buffer with Freund's incomplete adjuvant; and third injection, 100 μ g of protein in 0.5 ml buffer with Freund's incomplete

Blood containing the antibodies was collected 70 days after the first injection. The antibodies were recovered and successfully used in an enzyme linked

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immunoabsorbent assay to detect the presence of virus particles (i.e., coat protein) of RSPaV-1 from a variety of tissue types of grapevines infected with RSP.

The antibodies produced against the expressed RSPaV-1 coat protein, therefore, are useful in the identification of the particles associated with RSP disease of grapevines, in the purification of the particles of RSPaV-1, and in the development of a serological diagnosis for RSP in grapevine. The use of the antibodies is suitable for detecting different strains of RSPaV-1. Because the coat proteins for strains RSP47-4 and RSP158 have high amino acid identities with the coat protein of RSPaV-1, it is very likely that the antibodies raised against RSPaV-1 coat protein will also detect other strains. Antibodies can be used in an ELISA to assay rapidly a large number of samples, thus making commercial development and utilization of diagnostic kits possible.

Example -19 Transformation of Grapevines with a Vector Containing RSPaV-1
Coat Protein Gene and Analysis of Transgenic Grapevines for
Resistance to RSP

The DNA molecule coding for the RSPaV-1 coat protein (e.g., SEQ. ID. No. 10) was cloned into a pEPT8 plant expression vector that contains the double 35S enhancer at restriction sites SalI and BamHI. The resulting recombinant plasmid, designated pEPT8/RSPaV-1 coat protein, was then cloned into the plant transformation vector pGA482G, which has resistance genes to gentamycin and tetracycline as selection markers. The resultant pGA482G containing pEPT8/RSPaV-1CP was used to transform grapevines using the *Agrobacterium* method.

The rootstock *Vitis rupestris* Scheele St. George was used in genetic transformation. Anthers were excised aseptically from flower buds. The pollen was crushed on a microscope slide with acetocarmine to observe the cytological stage (Bouquet et al., "Influence du Gentype sur la Production de cals: Dembryoides et Plantes Entieres par Culture Danthers in vitro dans le Genre Vitis," <u>C.R. Acad. Sci. Paris III</u> 295:560-74 (1982), which is hereby incorporated by reference). This was done to determine which stage was most favorable for callus induction.

Anthers were plated under aseptic condition at a density of 40 to 50 per 9 cm diameter Petri dish containing MSE. Plates were cultured at 28°C in the dark.

After 60 days, embryos were induced and transferred to hormone-free medium

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(HMG) for differentiation. Torpedo stage embryos were transferred to MGC medium to promote embryo germination. Cultures were maintained in the dark at 26-28°C and transferred to fresh medium at 3-4 week intervals. Elongated embryos were transferred to rooting medium (5-8 embryos per jar). The embryos were grown in a tissue culture room at 25°C with a daily 16 h photoperiod (76 μ mol. s) to induce shoot and root formation. After plants developed roots, they were transplanted to soil in the greenhouse.

The protocols used for transformation were modified from those described by Scorza et al., "Transformation of Grape (Vitis vinifera L.) Zygotic-Derived Somatic Embryos and Regeneration of Transgenic Plants," Plant Cell Rpt. 10 14:589-92 (1995), which is hereby incorporated by reference. Overnight cultures of Agrobacterium strain C58Z707 or LBA4404 were grown in LB medium at 28°C in a shaking incubator. Bacteria were centrifuged for 5 minutes at 3000-5000 rpm and resuspended in MS liquid medium (OD 1.0 at A600 nm). Calli with embryos were immersed in the bacterial suspension for 15-30 minutes, blotted dry, and transferred to 15 HMG medium with or without acetosyringone (100 μ M). Embryogenic calli were cocultivated with the bacteria for 48 h in the dark at 28°C. The plant material was then washed in MS liquid plus cefotaxime (300 mg/ml) and carbenicillin (200 mg/ml) 2-3 times. To select transgenic embryos, the material was transferred to HMG medium containing either 20 or 40 mg/L kanamycin, 300 mg/L cefotaxime, and 200 mg/L 20 carbenicillin. Alternatively, after co-cultivation, embryogenic calli were transferred to initiation MSE medium containing 25 mg/l kanamycin plus the same antibiotics listed above. All plant materials were incubated in continuous darkness at 28°C. After growth on selection medium for 3 months, embryos were transferred to HMG or MGC without kanamycin to promote elongation of embryos. They were then 25 transferred to rooting medium without antibiotics. Non-transformed calli were grown on the same media with and without kanamycin to verify the efficiency of the kanamycin selection process.

The X-gluc (5-bromo-4-chloro-3-indoyl-ß-glucuronidase)

histochemical assay was used to detect GUS (ß-glucuronidase) activity in embryos and plants that were transformed with constructs containing the GUS gene that survived kanamycin selection. All propagated plants were screened using an enzyme linked immunoabsorbent assay (ELISA) system (5 Prime-3 Prime, Boulder, Co.) to

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detect the NPTII (neomycin phosphotransferase II) protein in leaf extracts. ELISA tests with respective coat protein (CP)-specific antibodies were used to assay for CP. ELISA results were read on an SLT Spectra ELISA reader (Tecan U.S. Inc., Research Triangle Park, NC) 15-60 minutes after the substrate was added.

PCR analysis was carried out to detect the presence of transgene sequences in grape plants. Genomic DNA was isolated from transformed and non-transformed grape plants according to the method of Lodhi et al., "A Simple and Efficient Method for DNA Extraction from Grapevine Cultivars and Vitis Species," Plant Mol. Biol. Rpt. 12:6-13 (1994), which is hereby incorporated by reference.

Primer sets included those of specific primers to the transgene. DNA was initially denatured at 94°C for 3 minutes, then amplified by 35 cycles of 1 minute at 94°C (denaturing), 1 minute at 52°C (annealing), and 2 minutes at 72°C (polymerizing). Reaction samples were directly loaded and electrophoresed in 1.5 % agarose gels.

Southern analysis of transformants was accomplished by extracting genomic DNA from young leaves of transformed and non-transformed plants (3309C) as described above. DNA (10 µg) was digested with the restriction enzyme *Bgl* II, electrophoresed on a 0.8% agarose gel in TAE buffer and transferred to a Genescreen Plus membrane by capillary in 10 x SSC. A probe was prepared by random primer labeling of a PCR amplified gene coding sequence with radioisotope ³²P-dATP (Dupont, NEN). Pre-hybridization and hybridization steps were carried out at 65°C following the manufacturer's instruction. The autoradiograph was developed after overnight exposure.

Although the invention has been described in detail for the purposes of illustration, it is understood that such detail is solely for that purpose, and variations can be made therein by those skilled in the art without departing from the spirit and scope of the invention which is defined by the following claims.

SEQUENCE LISTING

(1) GENERAL INFORMATION:

- (i) APPLICANT: Cornell Research Foundation, Inc.
- (ii) TITLE OF INVENTION: RUPESTRIS STEM PITTING
 ASSOCIATED VIRUS NUCLIEC ACIDS,
 PROTEINS, AND THEIR USES
- (iii) NUMBER OF SEQUENCES: 54
 - (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: Nixon, Hargrave, Devans & Doyle LLP
 - (B) STREET: Clinton Square, P.O. Box 1051
 - (C) CITY: Rochester
 - (D) STATE: New York
 - (E) COUNTRY: U.S.A.
 - (F) ZIP: 14603
 - (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
 - (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER:
 - (B) FILING DATE:
 - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: US 60/047,147
 - (B) FILING DATE: 20-MAY-1997
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: US 60/069,902
 - (B) FILING DATE: 17-DEC-1997
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: Goldman, Michael L.
 - (B) REGISTRATION NUMBER: 30,727
 - (C) REFERENCE/DOCKET NUMBER: 19603/1723
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (716) 263-1304
 - (B) TELEFAX: (716) 263-1600

(2) INFORMATION FOR SEQ ID NO:1:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 8743 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

CGATAAACAT	AACAACAGAA	TCTGCATTGC	AGTAATATTC	CTTGAATATA	ATTGCAACGC	60
AATGGCCCTC	TCTTATAGGC	CTGCTGTTGA	AGAGGTGCTC	GCAAAATTCA	CCTCTGATGA	120
ACAATCCAGG	GTTTCTGCTA	CAGCTCTCAA	GGCATTAGTA	GACTTAGAGG	AAAGTCAGCA	180
CAATTTGTTC	TCTTTCGCAT	TGCCTGATAG	AAGCAAAGAA	AGGCTGATAT	CTTCTGGCAT	240
TTACTTAAGT	CCTTACAGTT	TCAGACCCCA	CTCACATCCA	GTTTGTAAAA	CTTTAGAAAA	300
TCACATTTTG	TACAATGTTT	TACCTAGTTA	TGTTAATAAT	TCATTTTACT	TTGTAGGAAT	360
CAAGGATTTT	AAGCTGCAGT	TCTTGAAAAG	GAGGAATAAG	GATCTCAGCT	TGGTAGCACT	420
CATAAATAGG	TTTGTGACAA	GTCGTGATGT	TAGTAGGTAT	GGGTCTGAGT	TCGTTATAAG	480
TTCTAGTGAC	AAATCAAGTC	AGGTTGTCAG	TAGAAAGGGC	ATTGGTGATT	CTAACACACT	540
CCGGAGATTG	GTCCCACGTG	TAATTTCCAC	AGGTGCCAGG	AATCTTTTTC	TGCATGATGA	600
GATTCACTAC	TGGTCAATTA	GTGATCTGAT	CAATTTTTTG	GACGTTGCCA	AGCCAAGCAT	660
GCTCTTGGCA	ACTGCAGTAA	TCCCTCCAGA	AGTGCTGGTT	GGCTCTCCAG	AGAGTCTTAA	720
CCCTTGGGCC	TACCAGTATA	AAATCAATGG	CAACCAACTG	CTCTTCGCAC	CAGATGGCAA	780
CTGGAATGAG	ATGTACTCAC	AACCTTTGTC	ATGCAGATAC	CTGCTCAAGG	CCAGATCTGT	840
AGTTCTGCCC	GATGGCTCAC	GCTACTCGGT	TGACATCATT	CACTCAAAAT	TTAGTCACCA	900
CTTGCTTAGT	TTCACCCCTA	TGGGTAATCT	TTTGACTTCA	AACATGCGAT	GTTTTTCTGG	960
CTTCGATGCA	ATAGGCATAA	AAGATCTTGA	ACCTCTAAGC	CGCGGCATGC	ACAGTTGCTT	1020
CCCAGTACAT	CATGATGTTG	TAACTAAGAT	ATATCTTTAT	TTGAGAACTC	TCAAGAAGCC	108
AGATAAGGAG	TCTGCCGAGG	CAAAGCTTCG	ACAACTCATA	GAAAAACCCA	CAGGGAGGGA	114
GATAAAGTTT	ATCGAGGATT	TTTCCTCACT	AGTAATAAAT	TGTGGGAGGA	GTGGCTCTTT	120
GCTTATGCCC	AACATTTCTA	AGTTGGTCAT	ATCATTCTTT	TGCCGGATGA	TGCCAAATGC	126
ACTOGOCAGG	ር ጥርጥርጥጥርጥ አ	CCTTTCCACA	に ずにですない。	C	. mcma.cmca.cm	120

TGAGCCCTTT	AATTTTTCCG	TTAATTTAGT	GGATATAACT	CCTGATTTCT	TTGAGCATTT	1380
ATTTCTCTTC	TCCTGCCTAA	ATGAGTTGAT	CGAGGAGGAC	GTTGAAGAGG	TCATGGACA_A	1440
TTCTTGGTTT	GGACTTGGGG	ACTTACAATT	CAATCGCCAG	AGGGCCCCGT	TCTTTCTTGG	1500
GTCTTCATAT	TGGCTCAACT	CCAAATTTTC	AGTTGAGCAC	AAGTTTTCAG	GCACCATCAA	1560
TTCTCAAATC	ATGCAAGTTA	TTTTATCTTT	GATCCCATTT	TCTGATGATC	CCACTTTTAG	1620
GCCATCTTCT	ACAGAGGTTA	ACCTTGCACT	ATCAGAGGTT	AAGGCTGCGC	TAGAAGCTAC	1680
TGGGCAGTCA	AAATTGTTCA	GGTTTTTGGT	GGACGACTGT	GCTATGCGTG	AGGTTAGAAG	1740
TTCCTATAAG	GTGGGCCTTT	TTAAGCACAT	AAAAGCCCTC	ACTCATTGCT	TTAATTCTTG	1800
TGGCCTCCAA	TGGTTCCTCC	TTAGGCAAAG	GTCCAACCTC	AAATTTCTGA	AGGACAGGGC	1860
ATCGTCCTTT	GCTGATCTTG	ATTGTGAGGT	TATCAAAGTT	TATCAGCTTG	TAACATCACA	1920
GGCAATACTT	CCTGAGGCTC	TGCTTAGCTT	GACCAAAGTC	TTTGTCAGGG	ATTCTGACTC	1980
AAAGGGTGTT	TCCATTCCCA	GATTGGTCTC	GAGAAATGAG	CTAGAGGAAC	TAGCTCACCC	2040
AGCTAATTCA	A GCCCTTGAGG	AGCCTCAATC	: AGTTGATTGT	AATGCAGGCA	GGGTTCAAGC	2100
AAGCGTTTC	A AGTTCCCAGO	AGCTTGCCG	CACCCACTCT	CTTGGTAGC	TTAAGTCATC	2160
AATTGAGACA	A GCTAACAAG	CTTTTAACT1	GGAGGAGCTA	AGGATCATG	A TTAGAGTCTT	2220
GCCGGAGGA	r TTTAACTGG	TGGCGAAGA	A CATTGGTTT	T AAAGACAGG	TGAGAGGCAG	2280
GGGTGCATC	A TTCTTCTCA	A AACCAGGAA	T TTCATGTCAT	r AGTTACAAT	G GTGGGAGCCA	2340
CACAAGCTT	A GGGTGGCCA	A AGTTCATGG	A TCAGATTCT	A AGCTCCACT	G GTGGACGTAA	2400
TTACTACAA	T TCATGCCTG	G CTCAGATCT	A TGAGGAAAA	r TCAAAATTG	G CTCTTCATAA	2460
GGATGATGA	G AGTTGCTAT	G AAATTGGGC	A CAAAGTTTT	G ACTGTTAAT	T TAATCGGCTC	2520
AGCAACTTT	C ACTATTAGT.	A AGTCGCGAA	A TTTGGTTGG	G GGTAATCAT	T GCAGCCTGAC	2580
AATTGGGCC	A AATGAGTTT	T TCGAAATGC	C TAGGGGCAT	G CAATGCAAT	T ACTTCCATGG	2640
GGTTTCCAA	T TGTACGCCA	G GGCGGGTAT	C GCTGACCTT	T AGGCGCCAA	A AGTTGGAAGA	2700
TGATGATTT	G ATCTTCATA	A ATCCACAGG	T GCCCATTGA	G CTCAATCAT	G AAAAGCTTGA	2760
CCGAAGTAT	G TGGCAGATG	G GCCTTCATO	G AATTAAGAA	A TCTATTTCT	TA TGAATGGCAC	2820
GAGTTTTAC	C TCAGACCTA	T GCTCTTGTT	T CTCTTGCCA	C AACTTTCAT	A AATTCAAGGA	2880
TCTCATCA	AT AACTTGAGA	T TGGCCCTAG	G AGCACAAGG	G CTAGGTCAG	ST GTGACAGGGT	2940
TGTGTTTGC	CA ACAACAGGI	C CTGGTCTAT	C TAAGGTTT	TA GAAATGCC	C GGAGCAAAAA	3000
GCAATCAAT	TT TTGGTTCT	G AAGGTGCC	CT ATCCATAGA	AA ACAGATTA'	IG GTCCAAAAGT	3060
CCTGGGGT	CT TTTGAAGT	TT TCAAAGGG	GA CTTTCACA	TT AAGAAGAT	GG AGGAAGGTTC	3120

AATTTTTGTA	ATAACGTACA	AGGCCCCAAT	TAGATCCACT	GGCAGGTTGA	GGGTTCACAG	3180
TTCAGAATGC	TCATTTTCCG	GATCCAAAGA	GGTATTGCTA	GGCTGCCAGA	TTGAGGCATG	3240
TGCTGATTAT	GATATTGATG	ATTTTAACAC	TTTCTCTGTG	CCTGGTGATG	GCAATTGCTT	3300
TTGGCATTCT	GTTGGTTTTT	TACTTAGCAC	TGATGGACTT	GCCCTAAAGG	CCGGTATTCG	3360
ATCTTTCGTG	GAGAGTGAGC	GCTTGGTAAG	TCCAGATCTT	TCAGCCCCAG	CAATTTCTAA	3420
ACAATTGGAA	GAGAATGCTT	ATGCCGAGAA	TGAGATGATC	GCATTATTCT	GCATTCGGCA	3480
CCACGTAAGG	CCTATAGTGA	TCACACCAGA	ATATGAAGTT	AGTTGGAAAT	TCGGGGAAGG	3540
TGAGTGGCCC	CTATGTGGAA	TTCTTTGCCT	TAAATCAAAT	CACTTCCAAC	CATGCGCCCC	3600
ACTGAATGGT	TGCATGATCA	CAGCCATTGC	TTCAGCACTT	GGAAGGCGTG	AAGTTGATGT	3660
GTTAAATTAT	CTGTGTAGAC	CCAGCACTAA	TCATATTTTT	GAGGAGCTTT	GTCAGGGAGG	3720
GGGCCTTAAC	ATGATGTATT	TAGCTGAAGC	TTTTGAGGCC	TTTGACATTT	GCGCTAAATG	3780
TGATATAAAT	GGAGAGATTG	AAGTGATTAA	TCCGTGTGGT	AAAATTTCTG	CATTGTTTGA	3840
CATAACTAAT	GAGCACATAA	GGCATGTTGA	GAAAATAGGT	AATGGCCCTC	AGAGCATAAA	3900
AGTGGATGAA	TTGCGGAAGG	TCAAGCGATC	CGCCCTCGAT	TTCCTTTCAA	TGAATGGGTC	3960
TAAAATAACC	TACTTCCCAA	GCTTTGAGCG	GGCTGAAAAG	TTGCAAGGAT	GTTTGCTAGG	4020
GGGCCTAACT	GGCGTTATAA	GTGATGAGAA	GTTCAGTGAT	GCAAAACCTT	GGCTTTCTGG	4080
TATATCTACT	ACTGATATTA	AGCCAAGGGA	ATTGACTGTC	GTGCTTGGTA	CATTTGGGGC	4140
TGGGAAGAGT	TTCTTGTACA	AGAGTTTCAT	GAAAAGGTCT	GAGGGTAAAT	TCGTAACCTT	4200
TGTTTCTCCC	AGACGTGCTT	TAGCAAATTC	AATCAAAAAT	GATCTTGAAA	TGGATGATAG	4260
CTGCAAAGTT	GCTAAAGCAG	GTAGGTCAAA	GAAGGAAGGG	TGGGATGTAG	TAACTTTTGA	4320
GGTTTTCCTT	AGAAAAGTTG	CAGGATTGAA	GGCTGGCCAC	TGTGTGATTT	TTGATGAGGT	4380
CCAGTTGTTT	CCTCCTGGAT	ACATCGATCT	ATGCTTGCTT	ATTATACGTA	GTGATGCTTT	4440
CATTTCACTT	GCTGGTGATC	CATGTCAAAG	CACATATGAC	TCGCAAAAGG	ATCGGGCAAT	4500
TTTGGGCGCT	GAGCAGAGTG	ACATACTTAG	ACTGCTTGAG	GGCAAAACGT	ATAGGTATAA	4560
CATAGAAAGC	AGGAGGTTTG	TGAACCCAAT	GTTCGAATCA	AGACTGCCAT	GTCACTTCAA	4620
AAAGGGCTCG	ATGACTGCCG	CTTTCGCTGA	TTATGCAATC	TTCCATAATA	TGCATGACTT	4680
TCTCCTGGCG	AGGTCAAAAG	GTCCCTTGGA	TGCCGTTTTG	GTTTCCAGTT	TTGAGGAGAA	4740
AAAGATAGTC	CAGTCCTACT	TTGGAATGAA	ACAGCTCACA	CTCACATTTG	GTGAATCAAC	4800
TGGGTTGAAT	TTCAAAAATG	GGGGAATTCT	CATATCACAT	GATTCCTTTC	ACACAGATGA	4860
TCGGCGGTGG	CTTACTGCTT	TATCTCGCTT	CAGCCACAAT	TTGGATTTGG	TGAACATCAC	4920

AGGTCTGAGG GTGGAAAGTT TTCTCTCGCA CTTTGCTGGC AAACCCCTCT ACCATTTTTT	4980
ACAGCCAAA AGTGGGGAGA ATGTCATACG AGATTTGCTC CCAGGTGAGC CTAACTTCTT	5040
PAGTGGCTTT AACGTTAGCA TTGGAAAGAA TGAAGGTGTT AGGGAGGAGA AGTTATGTGG	5100
GACCCATGG TTAAAAGTTA TGCTTTTCCT GGGTCAAGAT GAGGATTGTG AAGTTGAAGA	5160
SATGGAGTCA GAATGCTCAA ATGAAGAATG GTTTAAAACC CACATCCCCT TGAGTAATCT	5220
GGAGTCAACC AGGGCCAGGT GGGTGGGTAA AATGGCCTTG AAAGAGTATC GGGAGGTGCG	5280
TTGTGGTTAT GAAATGACTC AACAATTCTT TGATGAGCAT AGGGGTGGAA CTGGTGAGCA	5340
ACTGAGCAAT GCATGTGAGA GGTTTGAAAG CATTTACCCA AGGCATAAAG GAAATGATTC	5400
AATAACCTTC CTCATGGCTG TCCGAAAGCG TCTCAAATTT TCGAAGCCCC AGGTTGAAGC	5460
TGCCAAACTG AGGCGGGCCA AACCATATGG GAAATTCTTA TTAGATTCTT TCCTATCCAA	5520
AATCCCATTG AAAGCCAGTC ATAATTCCAT CATGTTTCAT GAAGCGGTAC AGGAGTTTGA	5580
GGCGAAGAAG GCTAGTAAGA GTGCAGCAAC TATAGAGAAT CATGCAGGTA GGTCATGCAG	5640
GGATTGGTTA TTAGATGTTG CTCTGATTTT TATGAAGTCA CAACACTGTA CTAAATTTGA	5700
CAACAGGCTT AGAGTAGCTA AAGCTGGGCA AACCCTTGCT TGCTTCCAAC ATGCTGTTCT	5760
GGTTCGCTTT GCACCCTATA TGAGATACAT TGAGAAAAAG CTAATGCAAG CTCTGAAGCC	5820
TAACTTCTAC ATCCATTCAG GGAAAGGTCT GACGAGCTGA ACGAGTGGGT CAGAACTAGA	5880
GGATTCACTG GAATTTGCAC AGAATCAGAC TACGAAGCCT TTGATGCTTC CCAAGACCAC	5940
TTCATCCTAG CATTCGAATT GCAGATAATG AAATTTTTGG GGTTACCTGA AGATTTAATT	6000
TTGGACTATG AATTCATAAA AATTCATTTG GGATCAAAGC TCGGATCATT CTCTATAATG	6060
AGGTTTACTG GGGAGGCCAG CACATTTCTG TTTAACACTA TGGCTAACAT GTTGTTCACC	6120
TTTCTGAGGT ACGAACTAAC AGGCTCTGAG TCAATAGCAT TTGCAGGTGA TGACATGTGT	6180
GCTAATCGAA GGTTGCGGCT TAAAACAGAG CATGAGGGTT TTCTGAACAT GATTTGCCTT	6240
AAGGCCAAGG TTCAGTTTGT TTCCAATCCC ACATTCTGCG GATGGTGTTT ATTTAAGGAA	6300
GGGATCTTCA AGAAGCCTCA ATTAATCTGG GAGCGGATAT GCATTGCTAG GGAGATGGGC	6360
AACCTGGAGA ATTGTATTGA CAATTATGCG ATAGAGGTCT CCTATGCATA CCGACTGGGA	6420
GAGCTAGCCA TTGAAATGAT GACCGAGGAA GAAGTGGAGG CCCATTATAA TTGTGTTAGA	6480
TTCTTGGTCA GGAACAAGCA TAAGATGAGA TGCTCAATTT CAGGCCTATT TGAAGCTATT	6540
GATTAGGCCT TAAGTATTTG GCATTATTTG AGTATTATGA ATAATTTAGT TAAAGCATTG	6600
TCAGCATTTG AGTTTGTAGG TGTTTTCAGT GTGCTTAAAT TTCCAGTAGT CATTCATAGT	6660
GTGCCTGGTA GTGGTAAAAG TAGTTTAATA AGGGAGCTAA TTTCCGAGGA TGAGAATTTC	6720

ATAGCTTTCA	CAGCAGGTGT	TCCAGACAGC	CCTAATCTCA	CAGGAAGGTA	CATTAAGCCT	6780
TATTCTCCAG	GGTGTGCAGT	GCCAGGGAAA	GTTAATATAC	TTGATGAGTA	CTTGTCCGTC	6840
CAAGATTTTT	CAGGTTTTGA	TGTGCTGTTC	TCGGACCCAT	ACCAAAACAT	CAGCATTCCT	6900
AAAGAGGCAC	ATTTCATCAA	GTCAAAAACT	TGTAGGTTTG	GCGTGAATAC	TTGCAAATAT	6960
CTTTCCTCCT	TCGGTTTTAA	GGTTAGCAGT	GACGGTTTGG	ACAAAGTCAT	TGTGGGGTCG	7020
CCTTTTACAC	TAGATGTTGA	AGGGGTGCTA	ATATGCTTTG	GTAAGGAGGC	AGTGGATCTC	7080
GCTGTTGCGC	ACAACTCTGA	ATTCAAATTA	CCTTGTGAAG	TTAGAGGTTC	AACTTTTAAC	7140
GTCGTAACTC	TTTTGAAATC	AAGAGATCCA	ACCCCAGAGG	ATAGGCACTG	GTTTTACATT	7200
GCTGCTACAA	GACACAGGGA	GAAATTGATA	ATCATGCAGT	AAGATGCCTT	TTCAGCAGCC	7260
TGCGAATTGG	GCAAAAACCA	TAACTCCATT	GACAGTTGGC	TTGGGCATTG	GGCTTGTGCT	7320
GCATTTTCTG	AGGAAGTCAA	ATCTACCTTA	TTCAGGGGAC	AACATCCATC	AATTCCCTCA	7380
CGGTGGGCGT	TACAGGGACG	GTACAAAAAG	TATAACTTAC	TGTGGTCCAA	AGCAATCCTT	7440
CCCCAGCTCT	GGGATATTCG	GCCAATCTGA	GAATTTTGTG	CCCTTAATGC	TTGTCATAGG	7500
TCTAATCGCA	TTCATACATG	TATTGTCTGT	TTGGAATTCT	GGTCTTGGTA	GGAATTGTAA	7560
TTGCCATCCA	AATCCTTGCT	CATGTAGACA	GCAGTAGTGG	CAACCACCAA	GGTTGCTTCA	7620
TTAGGGCCAC	TGGAGAGTCA	ATTTTGATTG	AAAACTGCGG	CCCAAGTGAG	GCCCTTGCAT	7680
CCACTGTGAA	GGAGGTGCTG	GGAGGTTTGA	AGGCTTTAGG	GGTTAGCCGT	GCTGTTGAAG	7740
AAATTGATTA	TCATTGTTAA	ATTGGCTGAA	TGGCAAGTCA	AATTGGGAAA	CTCCCCGGTG	7800
AATCAAATGA	GGCTTTTGAA	GCCCGGCTAA	AATCGCTGGA	GTTAGCTAGA	GCTCAAAAGC	7860
AGCCGGAAGG	TTCTAATGCA	CCACCTACTC	TCAGTGGCAT	TCTTGCCAAA	CGCAAGAGGA	7920
TTATAGAGAA	TGCACTTTCA	AAGACGGTGG	ACATGAGGGA	GGTTTTGAAA	CACGAAACGG	7980
TGGTGATTTC	CCCAAATGTC	ATGGATGAAG	GTGCAATAGA	CGAGCTGATI	CGTGCATTTG	8040
GTGAATCTGG	CATAGCTGAA	AGCGTGCAAT	TTGATGTGGC	CATAGATATA	GCACGTCACT	8100
GCTCTGATGT	TGGTAGCTCC	CAGAGGTCAA	CCCTGATTGG	CAAGAGTCCA	TTTTGTGACC	8160
TAAACAGATC	: AGAAATAGCT	GGGATTATAA	GGGAGGTGAC	CACATTACGT	AGATTTTGCA	8220
TGTACTATGO	: AAAAATCGTG	TGGAACATCC	ATCTGGAGAC	GGGGATACCA	CCAGCTAACT	8280
GGGCCAAGAA	AGGATTTAAT	GAGAATGAAA	AGTTTGCAGC	CTTTGATTT	TTCTTGGGAG	8340
TCACAGATGA	GAGTGCGCTT	GAACCAAAGG	GTGGAATTA	AAGAGCTCC	A ACGAAAGCTG	8400
AGATGGTTGC	TAATATCGCC	TCTTTTGAGG	TTCAAGTGC1	CAGACAAGC	T ATGGCTGAAG	8460
GCAAGCGGAG	TTCCAACCTT	GGAGAGATT	A GTGGTGGAAC	GGCTGGTGC	A CTCATCAACA	8520

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ACCCCTTTTC	AAATGTTACA	CATGAATGAG	GATGACGAAG	TCAGCGACAA	TTCCGCAGTC	8580
CAATAATTCC	CCGATTTCAA	GGCTGGGTTA	AGCCTGTTCG	CTGGAATACC	GTACTAATAG	8640
TATTCCCTTT	CCATGCTAAA	TCCTATTTAA	TATATAAGGT	GTGGAAAGTA	AAAGAAGATT	8700
TGGTGTGTTT	TTATAGTTTT	CATTCAAAAA	ААААААААА	AAA		8743

(2) INFORMATION FOR SEQ ID NO:2:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 6485 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

ATGGCCCTCT	CTTATAGGCC	TGCTGTTGAA	GAGGTGCTCG	CAAAATTCAC	CTCTGATGAA	60
CAATCCAGGG	TTTCTGCTAC	AGCTCTCAAG	GCATTAGTAG	ACTTAGAGGA	AAGTCAGCAC	120
AATTTGTTCT	CTTTCGCATT	GCCTGATAGA	AGCAAAGAAA	GGCTGATATC	TTCTGGCATT	180
TACTTAAGTC	CTTACAGTTT	CAGACCCCAC	TCACATCCAG	TTTGTAAAAC	TTTAGAAAAT	240
CACATTTTGT	ACAATGTTTT	ACCTAGTTAT	GTTAATAATT	CATTTTACTT	TGTAGGAATC	300
AAGGATTTTA	AGCTGCAGTT	CTTGAAAAGG	AGGAATAAGG	ATCTCAGCTT	GGTAGCACTC	360
ATAAATAGGT	TTGTGACAAG	TCGTGATGTT	AGTAGGTATG	GGTCTGAGTT	CGTTATAAGT	420
TCTAGTGACA	AATCAAGTCA	GGTTGTCAGT	AGAAAGGGCA	TTGGTGATTC	TAACACACTC	480
CGGAGATTGG	TCCCACGTGT	AATTTCCACA	GGTGCCAGGA	ATCTTTTTCT	GCATGATGAG	540
ATTCACTACT	GGTCAATTAG	TGATCTGATC	AATTTTTTGG	ACGTTGCCAA	GCCAAGCATG	600
CTCTTGGCAA	CTGCAGTAAT	CCCTCCAGAA	GTGCTGGTTG	GCTCTCCAGA	GAGTCTTAAC	660
CCTTGGGCCT	ACCAGTATAA	ĂATCAATGG C	AACCAACTGC	TCTTCGCACC	AGATGGCAAC	720
TGGAATGAGA	TGTACTCACA	ACCTTTGTCA	TGCAGATACC	TGCTCAAGGC	CAGATCTGTA	780
GTTCTGCCCG	ATGGCTCACG	CTACTCGGTT	GACATCATTC	ACTCAAAATT	TAGTCACCAC	840
TTGCTTAGTT	TCACCCTAT	GGGTAATCTT	TTGACTTCAA	ACATGCGATG	TTTTTCTGGC	900
TTCGATGCAA	TAGGCATAAA	AGATCTTGAA	CCTCTAAGCC	GCGGCATGCA	CAGTTGCTTC	960
CCAGTACATO	ATGATGTTGT	AACTAAGATA	A TATCTTTATT	TGAGAACTCT	CAAGAAGCCA	1020
GATAAGGAGT	CTGCCGAGGC	AAAGCTTCGA	CAACTCATAG	AAAAACCCAC	AGGGAGGAG	1080
ATAAAGTTTA	TCGAGGATTI	TTCCTCACTA	GTAATAAATI	GTGGGAGGAG	G TGGCTCTTTG	1140

CTTATGCCCA	ACATTTCTAA	GTTGGTCATA	TCATTCTTTT	GCCGGATGAT	GCCAAATGCA	1200
CTCGCCAGGC	TCTCTTCTAG	CTTTCGAGAG	TGTTCGCTAG	ATTCATTTGT	GTACTCACTT	1260
GAGCCCTTTA	ATTTTTCCGT	TAATTTAGTG	GATATAACTC	CTGATTTCTT	TGAGCATTTA	1320
TTTCTCTTCT	CCTGCCTAAA	TGAGTTGATC	GAGGAGGACG	TTGAAGAGGT	CATGGACAAT	1380
TCTTGGTTTG	GACTTGGGGA	CTTACAATTC	AATCGCCAGA	GGGCCCCGTT	CTTTCTTGGG	1440
TCTTCATATT	GGCTCAACTC	CAAATTTTCA	GTTGAGCACA	AGTTTTCAGG	CACCATCAAT	1500
TCTCAAATCA	TGCAAGTTAT	TTTATCTTTG	ATCCCATTTT	CTGATGATCC	CACTTTTAGG	1560
CCATCTTCTA	CAGAGGTTAA	CCTTGCACTA	TCAGAGGTTA	AGGCTGCGCT	AGAAGCTACT	1620
GGGCAGTCAA	AATTGTTCAG	GTTTTTGGTG	GACGACTGTG	CTATGCGTGA	GGTTAGAAGT	1680
TCCTATAAGG	TGGGCCTTTT	TAAGCACATA	AAAGCCCTCA	CTCATTGCTŢ	TAATTCTTGT	1740
GGCCTCCAAT	GGTTCCTCCT	TAGGCAAAGG	TCCAACCTCA	AATTTCTGAA	GGACAGGGCA	1800
TCGTCCTTTG	CTGATCTTGA	TTGTGAGGTT	ATCAAAGTTT	ATCAGCTTGT	AACATCACAG	1860
GCAATACTTC	CTGAGGCTCT	GCTTAGCTTG	ACCAAAGTCT	TTGTCAGGGA	TTCTGACTCA	1920
AAGGGTGTTT	CCATTCCCAG	ATTGGTCTCG	AGAAATGAGC	TAGAGGAACT	AGCTCACCCA	1980
GCTAATTCAG	CCCTTGAGGA	GCCTCAATCA	GTTGATTGTA	ATGCAGGCAG	GGTTCAAGCA	2040
AGCGTTTCAA	GTTCCCAGCA	GCTTGCCGAC	ACCCACTCTC	TTGGTAGCGT	TAAGTCATCA	2100
ATTGAGACAG	CTAACAAGGC	TTTTAACTTG	GAGGAGCTAA	GGATCATGAT	TAGAGTCTTG	2160
CCGGAGGATT	TTAACTGGGT	GGCGAAGAAC	ATTGGTTTT	AAGACAGGC	GAGAGGCAGG	2220
GGTGCATCAT	TCTTCTCAAA	ACCAGGAATT	TCATGTCATA	A GTTACAATGO	TGGGAGCCAC	2280
ACAAGCTTAG	GGTGGCCAAA	GTTCATGGAT	CAGATTCTA	A GCTCCACTG	G TGGACGTAAT	2340
TACTACAATT	CATGCCTGGC	TCAGATCTAT	GAGGAAAAT	CAAAATTGG	C TCTTCATAAG	2400
GATGATGAGA	A GTTGCTATG	AATTGGGCAG	AAAGTTTTG	A CTGTTAATT	T AATCGGCTCA	2460
GCAACTTTC	A CTATTAGTA	A GTCGCGAAAT	TTGGTTGGG	G GTAATCATT	G CAGCCTGACA	2520
ATTGGGCCA	A ATGAGTTTT	CGAAATGCC	r aggggcatg	C AATGCAATT	A CTTCCATGGG	2580
GTTTCCAAT	r GTACGCCAG	GCGGGTATC	G CTGACCTTT.	A GGCGCCAAA	A GTTGGAAGAT	2640
GATGATTTG	A TCTTCATAA	A TCCACAGGT	G CCCATTGAG	C TCAATCATG	A AAAGCTTGAC	2700
CGAAGTATG	r ggcagatgg	G CCTTCATGG	A ATTAAGAAA	т статттста	T GAATGGCACG	276
AGTTTTACC	r cagacctat	G CTCTTGTTT	C TCTTGCCAC	A ACTTTCATA	A ATTCAAGGAT	282
CTCATCAAT	A ACTTGAGAT	r ggccctagg	A GCACAAGGG	C TAGGTCAGT	G TGACAGGGTT	288
GTGTTTGCA	A CAACAGGTC	C TGGTCTATC	T AAGGTTTTA	G AAATGCCTC	G GAGCAAAAAG	294

CAATCAATTT TGGTTCTTGA AGGTGCCCTA TCCATAGAAA CAGATTATGG TCCAAAAGTC	3000
CTGGGGTCTT TTGAAGTTTT CAAAGGGGAC TTTCACATTA AGAAGATGGA GGAAGGTTCA	3060
ATTTTTGTAA TAACGTACAA GGCCCCAATT AGATCCACTG GCAGGTTGAG GGTTCACAGT	3120
TCAGAATGCT CATTTTCCGG ATCCAAAGAG GTATTGCTAG GCTGCCAGAT TGAGGCATGT	3180
GCTGATTATG ATATTGATGA TTTTAACACT TTCTCTGTGC CTGGTGATGG CAATTGCTTT	3240
TGGCATTCTG TTGGTTTTTT ACTTAGCACT GATGGACTTG CCCTAAAGGC CGGTATTCGA	3300
TCTTTCGTGG AGAGTGAGCG CTTGGTAAGT CCAGATCTTT CAGCCCCAGC AATTTCTAAA	3360
CAATTGGAAG AGAATGCTTA TGCCGAGAAT GAGATGATCG CATTATTCTG CATTCGGCAC	3420
CACGTAAGGC CTATAGTGAT CACACCAGAA TATGAAGTTA GTTGGAAATT CGGGGAAGGT	3480
GAGTGGCCCC TATGTGGAAT TCTTTGCCTT AAATCAAATC	3540
CTGAATGGTT GCATGATCAC AGCCATTGCT TCAGCACTTG GAAGGCGTGA AGTTGATGTG	3600
TTAAATTATC TGTGTAGACC CAGCACTAAT CATATTTTTG AGGAGCTTTG TCAGGGAGGG	3660
GGCCTTAACA TGATGTATTT AGCTGAAGCT TTTGAGGCCT TTGACATTTG CGCTAAATGT	3720
GATATAAATG GAGAGATTGA AGTGATTAAT CCGTGTGGTA AAATTTCTGC ATTGTTTGAC	3780
ATAACTAATG AGCACATAAG GCATGTTGAG AAAATAGGTA ATGGCCCTCA GAGCATAAAA	3840
GTGGATGAAT TGCGGAAGGT CAAGCGATCC GCCCTCGATT TCCTTTCAAT GAATGGGTCT	3900
AAAATAACCT ACTTCCCAAG CTTTGAGCGG GCTGAAAAGT TGCAAGGATG TTTGCTAGGG	3960
GGCCTAACTG GCGTTATAAG TGATGAGAAG TTCAGTGATG CAAAACCTTG GCTTTCTGGT	4020
ATATCTACTA CTGATATTAA GCCAAGGGAA TTGACTGTCG TGCTTGGTAC ATTTGGGGCT	4080
GGGAAGAGTT TCTTGTACAA GAGTTTCATG AAAAGGTCTG AGGGTAAATT CGTAACCTTT	4140
GTTTCTCCCA GACGTGCTTT AGCAAATTCA ATCAAAAATG ATCTTGAAAT GGATGATAGC	4200
TGCAAAGTTG CTAAAGCAGG TAGGTCAAAG AAGGAAGGGT GGGATGTAGT AACTTTTGAG	4260
GTTTTCCTTA GAAAAGTTGC AGGATTGAAG GCTGGCCACT GTGTGATTTT TGATGAGGTC	4320
CAGTTGTTTC CTCCTGGATA CATCGATCTA TGCTTGCTTA TTATACGTAG TGATGCTTTC	4380
ATTTCACTTG CTGGTGATCC ATGTCAAAGC ACATATGACT CGCAAAAGGA TCGGGCAATT	4440
TTGGGCGCTG AGCAGAGTGA CATACTTAGA CTGCTTGAGG GCAAAACGTA TAGGTATAAC	4500
ATAGAAAGCA GGAGGTTTGT GAACCCAATG TTCGAATCAA GACTGCCATG TCACTTCAAA	4560
AAGGGCTCGA TGACTGCCGC TTTCGCTGAT TATGCAATCT TCCATAATAT GCATGACTTT	4620
CTCCTGGCGA GGTCAAAAGG TCCCTTGGAT GCCGTTTTGG TTTCCAGTTT TGAGGAGAAA	4680
ANCHUNCTOR ACTOCTACTT TECANTENAN CACCTONON TENCATTEG TENATENACT	4740

GGGTTGAATT	TCAAAAATGG	GGGAATTCTC	ATATCACATG	ATTCCTTTCA	CACAGATGAT	4800
CGGCGGTGGC	TTACTGCTTT	ATCTCGCTTC	AGCCACAATT	TGGATTTGGT	GAACATCACA	4860
GGTCTGAGGG	TGGAAAGTTT	TCTCTCGCAC	TTTGCTGGCA	AACCCCTCTA	CCATTTTTA	4920
ACAGCCAAAA	GTGGGGAGAA	TGTCATACGA	GATTTGCTCC	CAGGTGAGCC	TAACTTCTTC	4980
AGTGGCTTTA	ACGTTAGCAT	TGGAAAGAAT	GAAGGTGTTA	GGGAGGAGAA	GTTATGTGGT	5040
GACCCATGGT	TAAAAGTTAT	GCTTTTCCTG	GGTCAAGATG	AGGATTGTGA	AGTTGAAGAG	5100
ATGGAGTCAG	AATGCTCAAA	TGAAGAATGG	TTTAAAACCC	ACATCCCCTT	GAGTAATCTG	5160
GAGTCAACCA	GGGCCAGGTG	GGTGGGTAAA	ATGGCCTTGA	AAGAGTATCG	GGAGGTGCGT	5220
TGTGGTTATG	AAATGACTCA	ACAATTCTTT	GATGAGCATA	GGGGTGGAAC	TGGTGAGCAA	5280
CTGAGCAATG	CATGTGAGAG	GTTTGAAAGC	ATTTACCCAA	GGCATAAAGG	AAATGATTCA	5340
ATAACCTTCC	TCATGGCTGT	CCGAAAGCGT	CTCAAATTTT	CGAAGCCCCA	GGTTGAAGCT	5400
GCCAAACTGA	GGCGGGCCAA	ACCATATGGG	AAATTCTTAT	TAGATTCTTT	CCTATCCAAA	5460
ATCCCATTGA	AAGCCAGTCA	TAATTCCATC	ATGTTTCATG	AAGCGGTACA	GGAGTTTGAG	5520
GCGAAGAAGG	CTAGTAAGAG	TGCAGCAACT	ATAGAGAATC	ATGCAGGTAG	GTCATGCAGG	5580
GATTGGTTAT	TAGATGTTGC	TCTGATTTTT	ATGAAGTCAC	AACACTGTAC	TAAATTTGAC	5640
AACAGGCTTA	GAGTAGCTAA	AGCTGGGCAA	ACCCTTGCTT	GCTTCCAACA	TGCTGTTCTG	5700
GTTCGCTTTG	CACCCTATAT	GAGATACATT	GAGAAAAAGC	TAATGCAAGC	TCTGAAGCCT	5760
AACTTCTACA	TCCATTCAGG	GAAAGGTCTG	ACGAGCTGAA	CGAGTGGGTC	AGAACTAGAG	5820
GATTCACTGG	AATTTGCACA	GAATCAGACT	ACGAAGCCTT	TGATGCTTCC	CAAGACCACT	5880
TCATCCTAGC	ATTCGAATTG	CAGATAATGA	AATTTTTGGG	GTTACCTGAA	GATTTAATTT	5940
TGGACTATGA	ATTCATAAAA	ATTCATTTGG	GATCAAAGCT	CGGATCATTC	TCTATAATGA	6000
GGTTTACTGG	GGAGGCCAGC	ACATTTCTGT	TTAACACTAT	GGCTAACATG	TTGTTCACCT	6060
TTCTGAGGTA	CGAACTAACA	GGCTCTGAGT	CAATAGCATT	TGCAGGTGAT	GACATGTGTG	6120
CTAATCGAAG	GTTGCGGCTT	AAAACAGAGC	ATGAGGGTTT	TCTGAACAT	ATTTGCCTTA	6180
AGGCCAAGGT	TCAGTTTGTT	TCCAATCCCA	CATTCTGCGG	ATGGTGTTT	TTTAAGGAAG	6240
GGATCTTCAA	GAAGCCTCAA	TTAATCTGGG	AGCGGATATG	CATTGCTAG	GAGATGGGCA	6300
ACCTGGAGAA	TTGTATTGAC	AATTATGCGA	TAGAGGTCTC	CTATGCATAC	CGACTGGGAG	6360
AGCTAGCCAT	TGAAATGATG	ACCGAGGAAG	AAGTGGAGGG	CCATTATAA	T TGTGTTAGAT	6420
TCTTGGTCAG	GAACAAGCAI	AAGATGAGAT	GCTCAATTT	AGGCCTATT	I GAAGCTATTG	6480
ATTAG						6485

(2) INFORMATION FOR SEQ ID NO:3:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 2161 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Met Ala Leu Ser Tyr Arg Pro Ala Val Glu Glu Val Leu Ala Lys Phe 1 5 10 15

Thr Ser Asp Glu Gln Ser Arg Val Ser Ala Thr Ala Leu Lys Ala Leu 20 25 30

Val Asp Leu Glu Glu Ser Gln His Asn Leu Phe Ser Phe Ala Leu Pro 35 40 45

Asp Arg Ser Lys Glu Arg Leu Ile Ser Ser Gly Ile Tyr Leu Ser Pro 50 55 60

Tyr Ser Phe Arg Pro His Ser His Pro Val Cys Lys Thr Leu Glu Asn 65 70 75 80

His Ile Leu Tyr Asn Val Leu Pro Ser Tyr Val Asn Asn Ser Phe Tyr 85 90 95

Phe Val Gly Ile Lys Asp Phe Lys Leu Gln Phe Leu Lys Arg Arg Asn 100 105 110

Lys Asp Leu Ser Leu Val Ala Leu Ile Asn Arg Phe Val Thr Ser Arg 115 120 125

Asp Val Ser Arg Tyr Gly Ser Glu Phe Val Ile Ser Ser Ser Asp Lys 130 135 140

Ser Ser Gln Val Val Ser Arg Lys Gly Ile Gly Asp Ser Asn Thr Leu 145 150 155 160

Arg Arg Leu Val Pro Arg Val Ile Ser Thr Gly Ala Arg Asn Leu Phe 165 170 175

Leu His Asp Glu Ile His Tyr Trp Ser Ile Ser Asp Leu Ile Asn Phe 180 185 190

Leu Asp Val Ala Lys Pro Ser Met Leu Leu Ala Thr Ala Val Ile Pro 195 200 205

Pro Glu Val Leu Val Gly Ser Pro Glu Ser Leu Asn Pro Trp Ala Tyr 210 215 220

Gln Tyr Lys Ile Asn Gly Asn Gln Leu Leu Phe Ala Pro Asp Gly Asn 225 230 235 240

Trp Asn Glu Met Tyr Ser Gln Pro Leu Ser Cys Arg Tyr Leu Leu Lys Ala Arg Ser Val Val Leu Pro Asp Gly Ser Arg Tyr Ser Val Asp Ile 260 265 Ile His Ser Lys Phe Ser His His Leu Leu Ser Phe Thr Pro Met Gly 280 Asn Leu Leu Thr Ser Asn Met Arg Cys Phe Ser Gly Phe Asp Ala Ile 290 300 295 Gly Ile Lys Asp Leu Glu Pro Leu Ser Arg Gly Met His Ser Cys Phe 310 315 Pro Val His His Asp Val Val Thr Lys Ile Tyr Leu Tyr Leu Arg Thr 325 330 Leu Lys Lys Pro Asp Lys Glu Ser Ala Glu Ala Lys Leu Arg Gln Leu Ile Glu Lys Pro Thr Gly Arg Glu Ile Lys Phe Ile Glu Asp Phe Ser Ser Leu Val Ile Asn Cys Gly Arg Ser Gly Ser Leu Leu Met Pro Asn Ile Ser Lys Leu Val Ile Ser Phe Phe Cys Arg Met Met Pro Asn Ala Leu Ala Arg Leu Ser Ser Phe Arg Glu Cys Ser Leu Asp Ser Phe Val Tyr Ser Leu Glu Pro Phe Asn Phe Ser Val Asn Leu Val Asp Ile Thr Pro Asp Phe Phe Glu His Leu Phe Leu Phe Ser Cys Leu Asn Glu Leu Ile Glu Glu Asp Val Glu Glu Val Met Asp Asn Ser Trp Phe Gly Leu Gly Asp Leu Gln Phe Asn Arg Gln Arg Ala Pro Phe Phe Leu Gly Ser Ser Tyr Trp Leu Asn Ser Lys Phe Ser Val Glu His Lys Phe Ser 490 Gly Thr Ile Asn Ser Gln Ile Met Gln Val Ile Leu Ser Leu Ile Pro Phe Ser Asp Asp Pro Thr Phe Arg Pro Ser Ser Thr Glu Val Asn Leu 520 Ala Leu Ser Glu Val Lys Ala Ala Leu Glu Ala Thr Gly Gln Ser Lys 530 535 Leu Phe Arg Phe Leu Val Asp Asp Cys Ala Met Arg Glu Val Arg Ser

550

555

Ser Tyr Lys Val Gly Leu Phe Lys His Ile Lys Ala Leu Thr His Cys 570 Phe Asn Ser Cys Gly Leu Gln Trp Phe Leu Leu Arg Gln Arg Ser Asn 585 Leu Lys Phe Leu Lys Asp Arg Ala Ser Ser Phe Ala Asp Leu Asp Cys Glu Val Ile Lys Val Tyr Gln Leu Val Thr Ser Gln Ala Ile Leu Pro 615 Glu Ala Leu Leu Ser Leu Thr Lys Val Phe Val Arg Asp Ser Asp Ser 630 635 625 Lys Gly Val Ser Ile Pro Arg Leu Val Ser Arg Asn Glu Leu Glu Glu 645 650 Leu Ala His Pro Ala Asn Ser Ala Leu Glu Glu Pro Gln Ser Val Asp 665 660 Cys Asn Ala Gly Arg Val Gln Ala Ser Val Ser Ser Ser Gln Gln Leu Ala Asp Thr His Ser Leu Gly Ser Val Lys Ser Ser Ile Glu Thr Ala Asn Lys Ala Phe Asn Leu Glu Glu Leu Arg Ile Met Ile Arg Val Leu Pro Glu Asp Phe Asn Trp Val Ala Lys Asn Ile Gly Phe Lys Asp Arg Leu Arg Gly Arg Gly Ala Ser Phe Phe Ser Lys Pro Gly Ile Ser Cys 745 His Ser Tyr Asn Gly Gly Ser His Thr Ser Leu Gly Trp Pro Lys Phe Met Asp Gln Ile Leu Ser Ser Thr Gly Gly Arg Asn Tyr Tyr Asn Ser 775 Cys Leu Ala Gln Ile Tyr Glu Glu Asn Ser Lys Leu Ala Leu His Lys 790 Asp Asp Glu Ser Cys Tyr Glu Ile Gly His Lys Val Leu Thr Val Asn Leu Ile Gly Ser Ala Thr Phe Thr Ile Ser Lys Ser Arg Asn Leu Val 820 825 Gly Gly Asn His Cys Ser Leu Thr Ile Gly Pro Asn Glu Phe Phe Glu 840 Met Pro Arg Gly Met Gln Cys Asn Tyr Phe His Gly Val Ser Asn Cys 855 Thr Pro Gly Arg Val Ser Leu Thr Phe Arg Arg Gln Lys Leu Glu Asp

870

875

- Asp Asp Leu Ile Phe Ile Asn Pro Gln Val Pro Ile Glu Leu Asn His 885 890 895
- Glu Lys Leu Asp Arg Ser Met Trp Gln Met Gly Leu His Gly Ile Lys 900 905 910
- Lys Ser Ile Ser Met Asn Gly Thr Ser Phe Thr Ser Asp Leu Cys Ser 915 920 925
- Cys Phe Ser Cys His Asn Phe His Lys Phe Lys Asp Leu Ile Asn Asn 930 935 940
- Leu Arg Leu Ala Leu Gly Ala Gln Gly Leu Gly Gln Cys Asp Arg Val 945 950 955 960
- Val Phe Ala Thr Thr Gly Pro Gly Leu Ser Lys Val Leu Glu Met Pro 965 970 975
- Arg Ser Lys Lys Gln Ser Ile Leu Val Leu Glu Gly Ala Leu Ser Ile 980 985 990
- Glu Thr Asp Tyr Gly Pro Lys Val Leu Gly Ser Phe Glu Val Phe Lys 995 1000 1005
- Gly Asp Phe His Ile Lys Lys Met Glu Glu Gly Ser Ile Phe Val Ile 1010 1015 1020
- Thr Tyr Lys Ala Pro Ile Arg Ser Thr Gly Arg Leu Arg Val His Ser 1025 1030 1035 1040
- Ser Glu Cys Ser Phe Ser Gly Ser Lys Glu Val Leu Leu Gly Cys Gln 1045 1050 1055
- Ile Glu Ala Cys Ala Asp Tyr Asp Ile Asp Asp Phe Asn Thr Phe Ser 1060 1065 1070
- Val Pro Gly Asp Gly Asn Cys Phe Trp His Ser Val Gly Phe Leu Leu 1075 1080 1085
- Ser Thr Asp Gly Leu Ala Leu Lys Ala Gly Ile Arg Ser Phe Val Glu 1090 1095 1100
- Ser Glu Arg Leu Val Ser Pro Asp Leu Ser Ala Pro Ala Ile Ser Lys 1105 1110 1115 1120
- Gln Leu Glu Glu Asn Ala Tyr Ala Glu Asn Glu Met Ile Ala Leu Phe 1125 1130 1135
- Cys Ile Arg His His Val Arg Pro Ile Val Ile Thr Pro Glu Tyr Glu 1140 1145 1150
- Val Ser Trp Lys Phe Gly Glu Gly Glu Trp Pro Leu Cys Gly Ile Leu 1155 1160 1165
- Cys Leu Lys Ser Asn His Phe Gln Pro Cys Ala Pro Leu Asn Gly Cys 1170 1175 1180
- Met Ile Thr Ala Ile Ala Ser Ala Leu Gly Arg Arg Glu Val Asp Val 1185 1190 1195 1200

- Leu Asn Tyr Leu Cys Arg Pro Ser Thr Asn His Ile Phe Glu Glu Leu 1205 1210 1215
- Cys Gln Gly Gly Leu Asn Met Met Tyr Leu Ala Glu Ala Phe Glu 1220 1225 1230
- Ala Phe Asp Ile Cys Ala Lys Cys Asp Ile Asn Gly Glu Ile Glu Val 1235 1240 1245
- Ile Asn Pro Cys Gly Lys Ile Ser Ala Leu Phe Asp Ile Thr Asn Glu 1250 1255 1260
- His Ile Arg His Val Glu Lys Ile Gly Asn Gly Pro Gln Ser Ile Lys 1265 1270 1275 1280
- Val Asp Glu Leu Arg Lys Val Lys Arg Ser Ala Leu Asp Phe Leu Ser 1285 1290 1295
- Met Asn Gly Ser Lys Ile Thr Tyr Phe Pro Ser Phe Glu Arg Ala Glu 1300 1305 1310
- Lys Leu Gln Gly Cys Leu Leu Gly Gly Leu Thr Gly Val Ile Ser Asp 1315 1320 1325
- Glu Lys Phe Ser Asp Ala Lys Pro Trp Leu Ser Gly Ile Ser Thr Thr 1330 1335 1340
- Asp Ile Lys Pro Arg Glu Leu Thr Val Val Leu Gly Thr Phe Gly Ala 1345 1350 1355 1360
- Gly Lys Ser Phe Leu Tyr Lys Ser Phe Met Lys Arg Ser Glu Gly Lys 1365 1370 1375
- Phe Val Thr Phe Val Ser Pro Arg Arg Ala Leu Ala Asn Ser Ile Lys 1380 1385 1390
- Asn Asp Leu Glu Met Asp Asp Ser Cys Lys Val Ala Lys Ala Gly Arg 1395 1400 1405
- Ser Lys Lys Glu Gly Trp Asp Val Val Thr Phe Glu Val Phe Leu Arg 1410 1415 1420
- Lys Val Ala Gly Leu Lys Ala Gly His Cys Val Ile Phe Asp Glu Val 1425 1430 1435 1446
- Gln Leu Phe Pro Pro Gly Tyr Ile Asp Leu Cys Leu Leu Ile Ile Arg 1445 1450 1455
- Ser Asp Ala Phe Ile Ser Leu Ala Gly Asp Pro Cys Gln Ser Thr Tyr 1460 1465 1470
- Asp Ser Gln Lys Asp Arg Ala Ile Leu Gly Ala Glu Gln Ser Asp Ile 1475 1480 1485
- Leu Arg Leu Leu Glu Gly Lys Thr Tyr Arg Tyr Asn Ile Glu Ser Arg 1490 1495 1500
- Arg Phe Val Asn Pro Met Phe Glu Ser Arg Leu Pro Cys His Phe Lys 1505 1510 1515 1520

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Lys Gly Ser Met Thr Ala Ala Phe Ala Asp Tyr Ala Ile Phe His Asn 1525 1530 1535

. Met His Asp Phe Leu Leu Ala Arg Ser Lys Gly Pro Leu Asp Ala Val 1540 1545 1550

Leu Val Ser Ser Phe Glu Glu Lys Lys Ile Val Gln Ser Tyr Phe Gly
1555 1560 1565

Met Lys Gln Leu Thr Leu Thr Phe Gly Glu Ser Thr Gly Leu Asn Phe 1570 1575 1580

Lys Asn Gly Gly Ile Leu Ile Ser His Asp Ser Phe His Thr Asp Asp 1585 1590 1595 1600

Arg Arg Trp Leu Thr Ala Leu Ser Arg Phe Ser His Asn Leu Asp Leu 1605 1610 1615

Val Asn Ile Thr Gly Leu Arg Val Glu Ser Phe Leu Ser His Phe Ala 1620 1625 1630

Gly Lys Pro Leu Tyr His Phe Leu Thr Ala Lys Ser Gly Glu Asn Val 1635 1640 1645

Ile Arg Asp Leu Leu Pro Gly Glu Pro Asn Phe Phe Ser Gly Phe Asn 1650 1655 1660

Val Ser Ile Gly Lys Asn Glu Gly Val Arg Glu Glu Lys Leu Cys Gly 1665 1670 1675 1680

Asp Pro Trp Leu Lys Val Met Leu Phe Leu Gly Gln Asp Glu Asp Cys 1685 1690 1695

Glu Val Glu Glu Met Glu Ser Glu Cys Ser Asn Glu Glu Trp Phe Lys 1700 1705 1710

Thr His Ile Pro Leu Ser Asn Leu Glu Ser Thr Arg Ala Arg Trp Val 1715 1720 1725

Gly Lys Met Ala Leu Lys Glu Tyr Arg Glu Val Arg Cys Gly Tyr Glu 1730 1735 1740

Met Thr Gln Gln Phe Phe Asp Glu His Arg Gly Gly Thr Gly Glu Gln 1745 1750 1755 1760

Leu Ser Asn Ala Cys Glu Arg Phe Glu Ser Ile Tyr Pro Arg His Lys 1765 1770 1775

Gly Asn Asp Ser Ile Thr Phe Leu Met Ala Val Arg Lys Arg Leu Lys 1780 1785 1790

Phe Ser Lys Pro Gln Val Glu Ala Ala Lys Leu Arg Arg Ala Lys Pro 1795 1800 1805

Tyr Gly Lys Phe Leu Leu Asp Ser Phe Leu Ser Lys Ile Pro Leu Lys 1810 1815 1820

Ala Ser His Asn Ser Ile Met Phe His Glu Ala Val Gln Glu Phe Glu 1825 1830 1835 1840

- Ala Lys Lys Ala Ser Lys Ser Ala Ala Thr Ile Glu Asn His Ala Gly
 1845 1850 1855
- Arg Ser Cys Arg Asp Trp Leu Leu Asp Val Ala Leu Ile Phe Met Lys 1860 1865 1870
 - Ser Gln His Cys Thr Lys Phe Asp Asn Arg Leu Arg Val Ala Lys Ala 1875 1880 1885
 - Gly Gln Thr Leu Ala Cys Phe Gln His Ala Val Leu Val Arg Phe Ala 1890 1895 1900
 - Pro Tyr Met Arg Tyr Ile Glu Lys Lys Leu Met Gln Ala Leu Lys Pro 1905 1910 1915 1920
 - Asn Phe Tyr Ile His Ser Gly Lys Gly Leu Asp Glu Leu Asn Glu Trp 1925 1930 1935
 - Val Arg Thr Arg Gly Phe Thr Gly Ile Cys Thr Glu Ser Asp Tyr Glu 1940 1945 1950
 - Ala Phe Asp Ala Ser Gln Asp His Phe Ile Leu Ala Phe Glu Leu Gln 1955 1960 1965
 - Ile Met Lys Phe Leu Gly Leu Pro Glu Asp Leu Ile Leu Asp Tyr Glu 1970 1975 1980
 - Phe Ile Lys Ile His Leu Gly Ser Lys Leu Gly Ser Phe Ser Ile Met 1985 1990 1995 2000
 - Arg Phe Thr Gly Glu Ala Ser Thr Phe Leu Phe Asn Thr Met Ala Asn 2005 2010 2015
 - Met Leu Phe Thr Phe Leu Arg Tyr Glu Leu Thr Gly Ser Glu Ser Ile 2020 2025 2030
 - Ala Phe Ala Gly Asp Asp Met Cys Ala Asn Arg Arg Leu Arg Leu Lys 2035 2040 2045
 - Thr Glu His Glu Gly Phe Leu Asn Met Ile Cys Leu Lys Ala Lys Val 2050 2055 2060
 - Gln Phe Val Ser Asn Pro Thr Phe Cys Gly Trp Cys Leu Phe Lys Glu 2065 2070 2075 2080
 - Gly Ile Phe Lys Lys Pro Gln Leu Ile Trp Glu Arg Ile Cys Ile Ala 2085 2090 2095
 - Arg Glu Met Gly Asn Leu Glu Asn Cys Ile Asp Asn Tyr Ala Ile Glu 2100 2105 2110
 - Val Ser Tyr Ala Tyr Arg Leu Gly Glu Leu Ala Ile Glu Met Met Thr 2115 2120 2125
 - Glu Glu Glu Val Glu Ala His Tyr Asn Cys Val Arg Phe Leu Val Arg 2130 2135 2140
 - Asn Lys His Lys Met Arg Cys Ser Ile Ser Gly Leu Phe Glu Ala Ile 2145 2150 2155 2160

Asp

(2)	INFORMATION	FOR	SEQ	ID	NO:4:
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- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 663 base pairs

 - (B) TYPE: nucleic acid(C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

ATGAATAATT	TAGTTAAAGC	ATTGTCAGCA	TTTGAGTTTG	TAGGTGTTTT	CAGTGTGCTT	60
AAATTTCCAG	TAGTCATTCA	TAGTGTGCCT	GGTAGTGGTA	AAAGTAGTTT	AATAAGGGAG	120
CTAATTTCCG	AGGATGAGAA	TTTCATAGCT	TTCACAGCAG	GTGTTCCAGA	CAGCCCTAAT	180
CTCACAGGAA	GGTACATTAA	GCCTTATTCT	CCAGGGTGTG	CAGTGCCAGG	GAAAGTTAAT	240
ATACTTGATG	AGTACTTGTC	CGTCCAAGAT	TTTTCAGGTT	TTGATGTGCT	GTTCTCGGAC	300
CCATACCAAA	ACATCAGCAT	TCCTAAAGAG	GCACATTTCA	TCAAGTCAAA	AACTTGTAGG	360
TTTGGCGTGA	ATACTTGCAA	ATATCTTTCC	TCCTTCGGTT	TTAAGGTTAG	CAGTGACGGT	420
TTGGACAAAG	TCATTGTGGG	GTCGCCTTTT	ACACTAGATG	TTGAAGGGGT	GCTAATATGC	480
TTTGGTAAGG	AGGCAGTGGA	TCTCGCTGTT	GCGCACAACT	CTGAATTCAA	ATTACCTTGT	540
GAAGTTAGAG	GTTCAACTTT	TAACGTCGTA	ACTCTTTTGA	AATCAAGAGA	TCCAACCCCA	600
GAGGATAGGC	ACTGGTTTTA	CATTGCTGCT	ACAAGACACA	GGGAGAAATT	GATAATCATG	660
CAG						663

(2) INFORMATION FOR SEQ ID NO:5:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 221 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Met Asn Asn Leu Val Lys Ala Leu Ser Ala Phe Glu Phe Val Gly Val

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Phe	Ser	Val	Leu 20	Lys	Phe	Pro	Val	Val 25	Ile	His	Ser	Val	Pro 30	Gly	Ser
Gly	Lys	Ser 35	Ser	Leu	Ile	Arg	Glu 40	Leu	Ile	Ser	Glu	Asp 45	Glu	Asn	Phe
Ile	Ala 50	Phe	Thr	Ala	Gly	Val 55	Pro	Asp	Ser	Pro	Asn 60	Leu	Thr	Gly	Arg
Tyr 65	Ile	Lys	Pro	Tyr	Ser 70	Pro	Gly	Суз	Ala	Val. 75	Pro	Gly	Lys	Val	Asn 80
Ile	Leu	Asp	Glu	Tyr 85	Leu	Ser	Val	Gln	Asp 90	Phe	Ser	Gly	Phe	Asp 95	Val
Leu	Phe	Ser	Asp 100	Pro	Tyr	Gln	Asn	Ile 105	Ser	Ile	Pro	Lys	Glu 110	Ala	His
Phe	Ile	Lys 115		Lys	Thr	Суѕ	Arg 120	Phe	Gly	Val	Asn	Thr 125		Lys	Tyr
Leu	Ser 130		Phe	Gly	Phe	Lys 135		Ser	Ser	Asp	Gly 140		Asp	Lys	Val
Ile 145		Gly	Ser	Pro	Phe 150		Leu	Asp	Val	Glu 155		Val	Leu	Ile	Cys 160
Phe	Gly	Lys	: Glu	Ala 165		Asp	Leu	Ala	Val		His	Asn	i Ser	Glu 175	Phe
Lys	Lev	Pro	Cys 180		ı Val	. Arg	g Gly	Ser 185		Phe	e Asr	val	Val		Lev
Lev	ı 'Lys	Ser 195		J Asp	Pro	Thi	200		ı Asp	Arq	g His	205		≘ Туг	: Ile
Ala	a Ala 210		r Ar	g His	s Arg	g Glu 21		s Let	ı Ile	e Ile	e Met		n		

(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 351 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

ATGCCTTTC AGCAGCCTGC GAATTGGGCA AAAACCATAA CTCCATTGAC AGTTGGCTTG 60

GGCATTGGGC TTGTGCTGCA TTTTCTGAGG AAGTCAAATC TACCTTATTC AGGGGACAAC 120

ATCCATCAAT TCCCTCACGG TGGGCGTTAC AGGGACGGTA CAAAAAGTAT AACTTACTGT 180

GGTCCAAAGC	AATCCTTCCC	CAGCTCTGGG	ATATTCGGCC	AATCTGAGAA	TTTTGTGCCC	•	240
TTAATGCTTG	TCATAGGTCT	AATCGCATTC	ATACATGTAT	TGTCTGTTTG	GAATTCTGGT		300
CTTGGTAGGA	ATTGTAATTG	CCATCCAAAT	CCTTGCTCAT	GTAGACAGCA	G		351

- (2) INFORMATION FOR SEQ ID NO:7:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 117 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

Met Pro Phe Gln Gln Pro Ala Asn Trp Ala Lys Thr Ile Thr Pro Leu
1 5 10 15

Thr Val Gly Leu Gly Ile Gly Leu Val Leu His Phe Leu Arg Lys Ser 20 25 30

Asn Leu Pro Tyr Ser Gly Asp Asn Ile His Gln Phe Pro His Gly Gly 35 40 45

Arg Tyr Arg Asp Gly Thr Lys Ser Ile Thr Tyr Cys Gly Pro Lys Gln 50 55 60

Ser Phe Pro Ser Ser Gly Ile Phe Gly Gln Ser Glu Asn Phe Val Pro 65 70 75 80

Leu Met Leu Val Ile Gly Leu Ile Ala Phe Ile His Val Leu Ser Val 85 90 95

Trp Asn Ser Gly Leu Gly Arg Asn Cys Asn Cys His Pro Asn Pro Cys
100 105 110

Ser Cys Arg Gln Gln 115

- (2) INFORMATION FOR SEQ ID NO:8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 240 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: cDNA
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

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60

120

ATGTATTGTC	TGTTTGGAAT	TCTGGTCTTG	GTAGGAATTG	TAATTGCCAT	CCAAATCCTT	60
GCTCATGTAG	ACAGCAGTAG	TGGCAACCAC	CAAGGTTGCT	TCATTAGGGC	CACTGGAGAG	120
TCAATTTTGA	TTGAAAACTG	CGGCCCAAGT	GAGGCCCTTG	CATCCACTGT	GAAGGAGGTG	180
CTCCCACCTT	TGAAGGCTTT	AGGGGTTAGC	CGTGCTGTTG	AAGAAATTGA	TTATCATTGT	240

(2) INFORMATION FOR SEQ ID NO:9:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 80 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

Met Tyr Cys Leu Phe Gly Ile Leu Val Leu Val Gly Ile Val Ile Ala 1 5 10 15

Ile Gln Ile Leu Ala His Val Asp Ser Ser Ser Gly Asn His Gln Gly 20 25 30

Cys Phe Ile Arg Ala Thr Gly Glu Ser Ile Leu Ile Glu Asn Cys Gly 35 40 45

Pro Ser Glu Ala Leu Ala Ser Thr Val Lys Glu Val Leu Gly Gly Leu 50 55 60

Lys Ala Leu Gly Val Ser Arg Ala Val Glu Glu Ile Asp Tyr His Cys 65 70 75 80

(2) INFORMATION FOR SEQ ID NO:10:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 777 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

ATGGCAAGTC AAATTGGGAA ACTCCCCGGT GAATCAAATG AGGCTTTTGA AGCCCGGCTA

AAATCGCTGG AGTTAGCTAG AGCTCAAAAG CAGCCGGAAG GTTCTAATGC ACCACCTACT

CTCAGT	GGCA	TTCTTGCCAA	ACGCAAGAGG	ATTATAGAGA	ATGCACTTTC	AAAGACGGTG	180
GACATG	AGGG	AGGTTTTGAA	ACACGAAACG	GTGGTGATTT	CCCCAAATGT	CATGGATGAA	240
GGTGCA	ATAG	ACGAGCTGAT	TCGTGCATTT	GGTGAATCTG	GCATAGCTGA	AAGCGTGCAA	300
TTTGAT	GTGG	CCATAGATAT	AGCACGTCAC	TGCTCTGATG	TTGGTAGCTC	CCAGAGTTCA	360
ACCCTG	ATTG	GCAAGAGTCC	ATTTTGTGAC	CTAAACAGAT	CAGAAATAGC	TGGGATTATA	420
AGGGAG	GTGA	CCACATTACG	TAGATTTTGC	ATGTACTATG	CAAAAATCGT	GTGGAACATC	480
CATCTG	GAGA	CGGGGATACC	ACCAGCTAAC	TGGGCCAAGA	AAGGATTTAA	TGAGAATGAA	540
AAGTTI	GCAG	CCTTTGATTT	TTTCTTGGGA	GTCACAGATG	AGAGTGCGCT	TGAACCAAAG	600
GGTGGA	ATTA	AAAGAGCTCC	AACGAAAGCT	GAGATGGTTG	CTAATATCGC	CTCTTTTGAG	660
GTTCAF	AGTGC	TCAGACAAGC	TATGGCTGAA	GGCAAGCGGA	GTTCCAACCT	TGGAGAGATT	72
AGTGGI	GGAA	CGGCTGGTGC	ACTCATCAAC	AACCCCTTTT	CAAATGTTAC	ACATGAA	77

(2) INFORMATION FOR SEQ ID NO:11:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 259 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

- Met Ala Ser Gln Ile Gly Lys Leu Pro Gly Glu Ser Asn Glu Ala Phe 1 5 10 15
- Glu Ala Arg Leu Lys Ser Leu Glu Leu Ala Arg Ala Gln Lys Gln Pro 20 25 30
- Glu Gly Ser Asn Ala Pro Pro Thr Leu Ser Gly Ile Leu Ala Lys Arg 35 40 45
- Lys Arg Ile Ile Glu Asn Ala Leu Ser Lys Thr Val Asp Met Arg Glu 50 55 60
- Val Leu Lys His Glu Thr Val Val Ile Ser Pro Asn Val Met Asp Glu 65 70 75 80
- Gly Ala Ile Asp Glu Leu Ile Arg Ala Phe Gly Glu Ser Gly Ile Ala 85 90 95
- Glu Ser Val Gln Phe Asp Val Ala Ile Asp Ile Ala Arg His Cys Ser 100 105 110
- Asp Val Gly Ser Ser Gln Ser Ser Thr Leu Ile Gly Lys Ser Pro Phe 115 120 125

255

Cys	Asp 130	Leu	Asn	Arg	Ser	Glu 135	Ile	Ala	Gly	Ile	Ile 140	Arg	Glu	Val	Thr	
Thr 145	Leu	Arg	Arg	Phe	Cys 150	Met	Tyr	Tyr	Ala	Lys 155	Ile	Val	Trp	Asn	Ile 160	
His	Leu	Glu	Thr	Gly 165	Ile	Pro	Pro	Ala	Asn 170	Trp	Ala	Lys	Lys	Gly 175	Phe	
Asn	Glu	Asn	Glu 180	Lys	Phe	Ala	Ala	Phe 185	Asp	Phe	Phe	Leu	Gly 190	Val	Thr	
Asp	Glu	Ser 195	Ala	Leu	Glu	Pro	Lys 200	Gly	Gly	Ile	Lys	Arg 205	Ala	Pro	Thr	
Lys	Ala 210		Met	Val	Ala	Asn 215	Ile	Ala	Ser	Phe	Glu 220	Val	Gln	Val	Leu	
Arg 225		Ala	Met	Ala	Glu 230		Lys	Arg	Ser	Ser 235	Asn	Leu	Gly	Glu	11e 240	
Ser	Gly	Gly	Thr	Ala	Gly	Ala	Lev	Ile	Asn	Asn	Pro	Phe	Ser	Asn	Val	

250

Thr His Glu

(2) INFORMATION FOR SEQ ID NO:12:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2680 base pairs(B) TYPE: nucleic acid
- (C) STRANDEDNESS: single

245

- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

GGCTGGGCAA	ACTTTGGCCT	GCTTTCAACA	CGCCGTCTTG	GTTCGCTTTG	CACCCTACAT	60
GCGATACATT	GAAAAGAAGC	TTGTGCAGGC	ATTGAAACCA	AATTTCTACA	TTCATTCTGG	120
CAAAGGTCTT	GATGAGCTAA	GTGAATGGGT	TAGAGCCAGA	GGTTTCACAG	GTGTGTGTAC	180
TGAGTCAGAC	TATGAAGCTT	TTGATGCATC	CCAAGATCAT	TTCATCCTGG	CATTTGAACT	240
GCAAATCATG	AGATTTTTAG	GACTGCCAGA	AGATCTGATT	TTAGATTATG	AGTTCATCAA	300
AATTCATCTT	GGGTCAAAGC	TTGGCTCTTT	TGCAATTATG	AGATTCACAG	GTGAGGCAAG	360
CACCTTCCTA	TTCAATACTA	TGGCCAACAT	GCTATTCACT	TTCCTGAGGT	ATGAGTTGAC	420
AGGTTCTGA	TCAATTGCAT	TTGCTGGAGA	TGATATGTGT	GCTAATCGCA	GGTTAAGACT	480
CAAGACTGAG	CACGCCGGCT	TTCTAAACAT	GATCTGTCTC	AAAGCTAAG	TGCAGTTTGT	540

1						
CACAAATCCC	ACCTTCTGTG	GATGGTGTTT	GTTTAAAGAG	GGAATCTTTA	AAAAACCCCA	600
GCTCATTTGG	GAAAGGATCT	GCATTGCTAG	GGAAATGGGT	AACTTGGACA	ATTGCATTGA	660
CAATTACGCA	ATTGAGGTGT	CTTATGCTTA	CAGACTTGGG	GAATTGTCCA	TAGGCGTGAT	720
GACTGAGGAG	GAAGTTGAAG	CACATTCTAA	CTGCGTGCGT	TTCCTGGTTC	GCAATAAGCA	780
CAAGATGAGG	TGCTCAATTT	CTGGTTTGTT	TGAAGTAATT	GTTTAGGCCT	TAAGTGTTTG	840
GCATGGTGTG	AGTATTATGA	ATAACTTAGT	CAAAGCTTTG	TCTGCTTTTG	AATTTGTTGG	900
TGTGTTTTGT	GTACTTAAAT	TTCCAGTTGT	TGTTCACAGT	GTTCCAGGTA	GCGGTAAAAG	960
TAGCCTAATA	AGGGAGCTCA	TTTCTGAAGA	CGAGGCTTTT	GTGGCCTTTA	CAGCAGGTGT	1020
GCCAGACAGT	CCAAATCTGA	CAGGGAGGTA	CATCAAGCCC	TACGCTCCAG	GGTGTGCAGT	1080
GCAAGGGAAA	ATAAACATAC	TTGATGAGTA	CTTGTCTGTC	TCTGATACTT	CTGGCTTTGA	1140
TGTGCTGTTC	TCAGACCCTT	ACCAGAATGT	CAGCATTCCA	AGGGAGGCAC	ACTTCATAAA	1200
AACCAAAACC	TGTAGGTTTG	GTACCAACAC	CTGCAAGTAC	CTTCAATCTT	TTGGCTTTAA	1260
TGTTTGTAGT	GATGGGGTGG	ATAAAGTTGT	TGTAGGGTCG	CCATTTGAAC	TGGAGGTTGA	1320
GGGGGTTCTC	ATTTGCTTTG	GAAAGGAGGC	TGTAGATCTA	GCAGTTGCAC	ACAATTCTGA	1380
CTTCAAGTTG	CCCTGCGAGG	TGCGGGGTTC	AACATTTGAC	GTTGTAACGT	TATTGAAGTC	1440
CAGGGATCCA	ACTTCAGAAG	ATAAGCATTG	GTTCTACGTT	GCAGCCACAA	GGCATCGAAG	1500
TAAACTGATA	ATAATGCAGT	AAAATGCCTT	TTCAGCAACC	TGCCAACTGG	GCTAAGACCA	1560
TAACTCCATT	AACTATTGGT	TTGGGCATTG	GGTTGGTTCT	GCACTTCTTA	AGGAAATCAA	1620
ATCTGCCATA	TTCAGGAGAC	AATATTCACC	AGTTCCCACA	CGGAGGGCAT	TACAGGGACG	1680
GCACGAAGAG	TATAACCTAT	TGTGGCCCTA	GGCAGTCATT	CCCAAGCTCA	GGAATATTCG	1740
GTCAGTCTGA	AAATTTCGTA	CCTCTAATAI	TGGTCGTGAC	TCTGGTCGCT	TTTATACATG	1800
CGTTATCTCT	TTGGAATTCT	GGTCCTAGT	GGAGTTGCAA	TTGCCATCCA	AATCCTTGCA	1860
CATGTAGACA	GCAGTAGTGG	CAACCATCA	A GGCTGTTTCA	TAAGAGCCAC	CGGGGAGTCA	1920
ATAGTAATTG	AGAATTGTGG	GCCGAGCGAC	GCCCTAGCTG	CTACAGTCA	A AGAGGTGTTG	1980
GGCGGTCTAA	AGGCTTTAGG	GGTTAGCCA	A AAGGTTGAT	AAATTAATT	A CAGTTGTTGA	2040
GACAGTTGAA	TGGCAAGTCA	AGTTGGAAA	A TTGCCTGGC	AATCAAATG	A AGCATATGAG	2100
GCTAGACTCA	AGGCTTTAGA	GTTAGCAAG	G GCCCAAAAA	G CTCCAGAAG	T CTCCAACCAA	2160
CCTCCCACAC	TTGGAGGCAT	TCTAGCCAA	A AGGAAAAGA	G TGATTGAGA	A TGCACTCTCA	2220
AAGACAGTGO	ATATGCGTG	A AGTCTTAAG	G CATGAATCT	G TTGTACTCT	C CCCGAATGTA	2280
ATGGACGAG	GAGCAATAG	A CGAGCTGAT	T CGTGCCTTT	G GGGAGTCGG	G CATAGCTGAA	2340

AATGTGCAGT	TTGATGTTGC	AATAGACATT	GCTCGCCACT	GTTCTGATGT	GGGGAGCTCT	2400
CAGAGGTCAA	CCCTTATTGG	TAAAAGCCCC	TTCTGTGAGT	TAAATAGGTC	TGAAATTGCC	2460
GGAATAATAA	GGGAGGTGAC	CACGCTGCGC	AGATTTTGCA	TGTACTACGC	AAAGATTGTG	2520
TGGAACATCC	ATTTGGAGAC	GGGAATACCA	CCAGCTAATT	GGGCCAAGAA	AGGATTTAAT	2580
GAGAATGAAA	AGTTTGCAGC	CTTTGACTTC	TTCCTTGGAG	TCACAGATGA	AAGCGCGCTT	2640
GAGCCTAAGG	GTGGAGTCAA	GAGAGCTCCA	ACAAAAGCAG			2680

(2) INFORMATION FOR SEQ ID NO:13:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 767 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

ATGCGATACA	TTGAAAAGAA	GCTTGTGCAG	GCATTGAAAC	CAAATTTCTA	CATTCATTCT	60
GGCAAAGGTC	TTGATGAGCT	AAGTGAATGG	GTTAGAGCCA	GAGGTTTCAC	AGGTGTGTGT	120
ACTGAGTCAG	ACTATGAAGC	TTTTGATGCA	TCCCAAGATC	ATTTCATCCT	GGCATTTGAA	180
CTGCAAATCA	TGAGATTTTT	AGGACTGCCA	GAAGATCTGA	TTTTAGATTA	TGAGTTCATC	240
AAAATTCATC	TTGGGTCAAA	GCTTGGCTCT	TTTGCAATTA	TGAGATTCAC	AGGTGAGGCA	300
AGCACCTTCC	TATTCAATAC	TATGGCCAAC	ATGCTATTCA	CTTTCCTGAG	GTATGAGTTG	360
ACAGGTTCTG	AATCAATTGC	ATTTGCTGGA	GATGATATGT	GTGCTAATCG	CAGGTTAAGA	420
CTCAAGACTG	AGCACGCCGG	CTTTCTAAAC	ATGATCTGTC	TCAAAGCTAA	GGTGCAGTTT	480
GTCACAAATC	CCACCTTCTG	TGGATGGTGT	TTGTTTAAAG	AGGGAATCTT	TAAAAAACCC	540
CAGCTCATTT	GGGAAAGGAT	CTGCATTGCT	AGGGAAATGG	GTAACTTGGA	CAATTGCATT	600
GACAATTACG	CAATTGAGGT	GTCTTATGCT	TACAGACTTG	GGGAATTGTC	CATAGGCGTG	660
ATGACTGAGG	AGGAAGTTGA	AGCACATTCT	AACTGCGTGC	GTTTCCTGGI	TCGCAATAAG	720
CACAAGATGA	GGTGCTCAAT	TTCTGGTTT	TTTGAAGTAA	TTGTTTA		767

(2) INFORMATION FOR SEQ ID NO:14:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 255 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:

(A)	LENGTH:	666	base	pairs
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- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:

ATGAATAACT	TAGTCAAAGC	TTTGTCTGCT	TTTGAATTTG	TTGGTGTGTT	TTGTGTACTT	60
AAATTTCCAG	TTGTTGTTCA	CAGTGTTCCA	GGTAGCGGTA	AAAGTAGCCT	AATAAGGGAG	120
CTCATTTCTG	AAGACGAGGC	TTTTGTGGCC	TTTACAGCAG	GTGTGCCAGA	CAGTCCAAAT	180
CTGACAGGGA	GGTACATCAA	GCCCTACGCT	CCAGGGTGTG	CAGTGCAAGG	GAAAATAAAC	240
ATACTTGATG	AGTACTTGTC	TGTCTCTGAT	ACTTCTGGCT	TTGATGTGCT	GTTCTCAGAC	300
CCTTACCAGA	ATGTCAGCAT	TCCAAGGGAG	GCACACTTCA	TAAAAACCAA	AACCTGTAGG	360
TTTGGTACCA	ACACCTGCAA	GTACCTTCAA	TCTTTTGGCT	TTAATGTTTG	TAGTGATGGG	420
GTGGATAAAG	TTGTTGTAGG	GTCGCCATTT	GAACTGGAGG	TTGAGGGGGT	TCTCATTTGC	48.0
TTTGGAAAGG	AGGCTGTAGA	TCTAGCAGTT	GCACACAATT	CTGACTTCAA	GTTGCCCTGC	540
GAGGTGCGGG	GTTCAACATT	TGACGTTGTA	ACGTTATTGA	AGTCCAGGGA	TCCAACTTCA	600
GAAGATAAGC	ATTGGTTCTA	CGTTGCAGCC	ACAAGGCATC	GAAGTAAACT	GATAATAATG	660
CAGTAA						666

(2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 221 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

Met Asn Asn Leu Val Lys Ala Leu Ser Ala Phe Glu Phe Val Gly Val 1 5 10 15

Phe Cys Val Leu Lys Phe Pro Val Val Val His Ser Val Pro Gly Ser 20 25 30

Gly Lys Ser Ser Leu Ile Arg Glu Leu Ile Ser Glu Asp Glu Ala Phe 35 40 45

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(2)	INFORMATION	FOR	SEO	ΙĐ	NO:18:
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- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 117 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

Met Pro Phe Gln Gln Pro Ala Asn Trp Ala Lys Thr Ile Thr Pro Leu
1 10 15

Thr Ile Gly Leu Gly Ile Gly Leu Val Leu His Phe Leu Arg Lys Ser 20 25 30

Asn Leu Pro Tyr Ser Gly Asp Asn Ile His Gln Phe Pro His Gly Gly 35 40 45

His Tyr Arg Asp Gly Thr Lys Ser Ile Thr Tyr Cys Gly Pro Arg Gln 50 55 60

Ser Phe Pro Ser Ser Gly Ile Phe Gly Gln Ser Glu Asn Phe Val Pro 65 70 75 80

Leu Ile Leu Val Val Thr Leu Val Ala Phe Ile His Ala Leu Ser Leu 85 90 95

Trp Asn Ser Gly Pro Ser Arg Ser Cys Asn Cys His Pro Asn Pro Cys 100 105 110

Thr Cys Arg Gln Gln 115

(2) INFORMATION FOR SEQ ID NO:19:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 243 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

ATGCGTTATC TCTTTGGAAT TCTGGTCCTA GTAGGAGTTG CAATTGCCAT CCAAATCCTT 60

GCACATGTAG ACAGCAGTAG TGGCAACCAT CAAGGCTGTT TCATAAGAGC CACCGGGGAG 120

TCAATAGTAA TTGAGAATTG TGGGCCGAGC GAGGCCCTAG CTGCTACAGT CAAAGAGGTG 180

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TTGGGCGGTC TAAAGGCTTT AGGGGTTAGC CAAAAGGTTG ATGAAATTAA TTACAGTTGT	240
TGA	243
(2) INFORMATION FOR SEQ ID NO:20:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 80 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: protein	-
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:	•
Met Arg Tyr Leu Phe Gly Ile Leu Val Leu Val Gly Val Ala Ile Ala 1 5 10 15	
Ile Gln Ile Leu Ala His Val Asp Ser Ser Ser Gly Asn His Gln Gly 20 25 30	
Cys Phe Ile Arg Ala Thr Gly Glu Ser Ile Val Ile Glu Asn Cys Gly 35 40 45	
Pro Ser Glu Ala Leu Ala Ala Thr Val Lys Glu Val Leu Gly Gly Leu 50 55 60	
Lys Ala Leu Gly Val Ser Gln Lys Val Asp Glu Ile Asn Tyr Ser Cys 65 70 75 80	
(2) INFORMATION FOR SEQ ID NO:21:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 631 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:	
ATGGCAAGTC AAGTTGGAAA ATTGCCTGGC GAATCAAATG AAGCATATGA GGCTAGACTC	60
AAGGCTTTAG AGTTAGCAAG GGCCCAAAAA GCTCCAGAAG TCTCCAACCA ACCTCCCACA	120
CTTGGAGGCA TTCTAGCCAA AAGGAAAAGA GTGATTGAGA ATGCACTCTC AAAGACAGTG	180
GATATGCGTG AAGTCTTAAG GCATGAATCT GTTGTACTCT CCCCGAATGT AATGGACGAG	240

GGAGCAATAG ACGAGCTGAT TCGTGCCTTT GGGGAGTCGG GCATAGCTGA AAATGTGCAG 300

175

TTTGATGTTG	CAATAGACAT	TGCTCGCCAC	TGTTCTGATG	TGGGGAGCTC	TCAGAGGTCA	360
ACCCTTATTG	GTAAAAGCCC	CTTCTGTGAG	TTAAATAGGT	CTGAAATTGC	ÇGGAATAATA	420
AGGGAGGTGA	CCACGCTGCG	CAGATTTTGC	ATGTACTACG	CAAAGATTGT	GTGGAACATC	480
CATTTGGAGA	CGGGAATACC	ACCAGCTAAT	TGGGCCAAGA	AAGGATTTAA	TGAGAATGAA	540
AAGTTTGCAG	CCTTTGACTT	CTTCCTTGGA	GTCACAGATG	AAAGCGCGCT	TGAGCCTAAG	606
GGTGGAGTCA	AGAGAGCTCC	AACAAAAGCA	G			63
		DO TO NO.22	_			

(2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 210 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

Met Ala Ser Gln Val Gly Lys Leu Pro Gly Glu Ser Asn Glu Ala Tyr 10 15 15 15 16 Glu Ala Arg Leu Lys Ala Leu Glu Leu Ala Arg Ala Gln Lys Ala Pro 20 25 30 Glu Val Ser Asn Gln Pro Pro Thr Leu Gly Gly Ile Leu Ala Lys Arg 45 Lys Arg Val Ile Glu Asn Ala Leu Ser Lys Thr Val Asp Met Arg Glu 50 60 Val Leu Arg His Glu Ser Val Val Leu Ser Pro Asn Val Met Asp Glu 65 70 70 75 Glu Asn Val Gln Phe Asp Val Ala Ile Asp Ile Ala Arg His Cys Ser 105 105 110 Asp Val Gly Ser Ser Gln Arg Ser Thr Leu Ile Gly Lys Ser Pro Phe 115 135 140 Thr Leu Arg Arg Phe Cys Met Tyr Tyr Ala Lys Ile Val Trp Asn Ile 145 155 160 Tyr 10 Tyr 10 Gly Glu Leu Arg Arg Phe Cys Met Tyr Tyr Ala Lys Ile Val Trp Asn Ile 145 150 160 Tyr 10 Tyr 10 Gly Glu Leu Arg Arg Phe Cys Met Tyr Tyr Ala Lys Ile Val Trp Asn Ile 145 150 160 Tyr 10 Tyr 10 Tyr Ala Lys Ile Val Trp Asn Ile 145 Tyr Asn Ile 155 160 Tyr Asn Val Cys Ile Val Trp Asn Ile 145 Tyr Tyr Ala Lys Ile Val Trp Asn Ile 145 Tyr Asn Ile 155 Tyr Asn Val Cys Ile Val Trp Asn Ile 145 Tyr Tyr Ala Lys Ile Val Trp Asn Ile 145 Tyr Tyr

His Leu Glu Thr Gly Ile Pro Pro Ala Asn Trp Ala Lys Lys Gly Phe

170

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Asn Glu Asn Glu Lys Phe Ala Ala Phe Asp Phe Phe Leu Gly Val Thr 185 190

Asp Glu Ser Ala Leu Glu Pro Lys Gly Gly Val Lys Arg Ala Pro Thr 200

Lys Ala 210

(2) INFORMATION FOR SEQ ID NO:23:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 2009 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

GAAGCTAGCA CATTTCTGTT CAACACTATG GCTAACATGT TGTTCACTTT TCTGAGATAT 60 GAACTGACGG GTTCAGAGTC AATAGCATTT GCAGGGGATG ATATGTGTGC TAATAGAAGG 120 TTGCGGCTTA AAACGGAGCA TGAGGGTTTT CTGAACATGA TCTGCCTTAA GGCCAAGGTT 180 CAGTTTGTTT CCAACCCCAC ATTCTGTGGA TGGTGCTTAT TTAAGGAGGG AATCTTCAAG 240 AAACCTCAAC TAATTTGGGA GCGAATATGC ATAGCCAGAG AGATGGGCAA TCTGGAGAAC 300 TGTATTGACA ATTATGCGAT AGAAGTGTCC TATGCATATA GATTGGGTGA GCTATCAATT 360 GAAATGATGA CAGAAGAAGA AGTGGAGGCA CACTACAATT GTGTGAGGTT CCTGGTTAGG 420 AACAAGCATA AGATGAGGTG CTCAATTTCA GGCCTGTTTG AAGTGGTTGA TTAGGCCTTA 480 AGTATTTGGC GTTGTTCGAG TTATTATGAA TAATTTAGTT AAAGCATTAT CAGCCTTCGA 540 GTTTATAGGT GTTTTCAATG TGCTCAAATT TCCAGTTGTT ATACATAGTG TGCCTGGTAG 600 TGGTAAGAGT AGCTTAATAA GGGAATTAAT CTCAGAGGAC GAGAGTTTCG TGGCTTTCAC 660 AGCAGGTGTT CCAGACAGTC CTAACCTCAC AGGGAGGTAC ATCAAGCCTT ACTCACCAGG 720 ATGCGCAGTG CAAGGAAAAG TGAATATACT TGATGAGTAC TTGTCCGTTC AAGACATTTC 780 GGGTTTTGAT GTACTGTTTT CAGACCCGTA CCAGAATATC AGTATTCCCC AAGAGGCGCA 840 TTTCATTAAG TCCAAGACTT GTAGGTTTGG TGTGAACACT TGCAAATACC TTTCCTCTTT 900 CGGTTTCGAA GTTAGCAGCG ACGGGCTGGA CGACGTCATT GTGGGATCGC CCTTCACTCT 960 AGATGTTGAA GGGGTGCTGA TATGTTTTGG CAAGGAGGCG GTAGATCTCG CTGTTGCGCA 1020 CAACTCTGAA TTCAAGTTGC CGTGTGAGGT TCGAGGTTCA ACCTTCAATG TGGTAACCCT

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TTGAAATCA	AGAGACCCAA	CCCCAGAGGA	CAGGCACTGG	TTTTACATCG	CTGCCACAAG	1140
ACATAGGAAG	AAATTGGTCÄ	TTATGCAGTA	AAATGCCTTT	TCAGCAGCCT	GCTAATTGGG	1200
CAAAAACCAT	AACTCCATTG	ACTATTGGCT	TAGGAATTGG	ACTTGTGCTG	CATTTTCTGA	1260
GAAAGTCAAA	TCTACCATAT	TCAGGAGACA	ACATCCATCA	ATTTCCTCAC	GGGGGGCGTT	1320
ACCGGGACGG	CACAAAAAGT	ATAACTTACT	GTGGCCCTAA	GCAGTCCTTC	CCCAGTTCAG	1380
GAATATTTGG	TCAGTCTGAG	AATTTTGTGC	CCTTAATGCT	TGTCATAGGT	CTAATTGCAT	1440
TCATACATGT	ATTGTCTGTT	TGGAATTCTG	GTCTTGGTAG	GAATTGCAAT	TGCCATCCAA	1500
ATCCTTGCTC	ATGTAGACAA	CAGTAGTGGC	AGTCACCAAG	GTTGCTTTAT	CAGGGCCACT	1560
GGAGAGTCTA	TTTTGATTGA	AAATTGTGGC	CCAAGCGAGG	CCCTTGCATC	AACAGTGAGG	1620
GAGGTGTTGG	GGGGTTTGAA	GGCTTTAGGA	ATTAGCCATA	CTACTGAAGA	AATTGATTAT	1680
CGTTGTTAAA	TTGGTTAAAT	GGCGAGTCAA	GTTGGTAAGC	TCCCCGGAGA	ATCAAATGAG	1740
GCATTTGAAG	CCCGGCTGAA	ATCACTGGAG	TTGGCTAGAG	CTCAAAAGCA	A GCCAGAAGGT	1800
TCAAACACAC	CGCCTACTCT	CAGTGGTGT	CTTGCCAAAC	GTAAGAGGG	TATTGAGAAT	1860
GCACTCTCAA	AGACAGTGGA	A CATGAGGGA	GTGTTGAAAC	ACGAAACGG	I TGTAATTTCC	1920
CCAAATGTCA	TGGATGAGG	TGCAATAGA	GAACTGATT	GTGCATTCG	G AGAATCAGGC	1980
ATAGCTGAGA	GCGCACAATT	TGATGTGGC				200

(2) INFORMATION FOR SEQ ID NO:24:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 447 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:

GAAGCTAGCA CATTTCTGTT CAACACTATG GCTAACATGT TGTTCACTTT TCTGAGATAT 60

GAACTGACGG GTTCAGAGTC AATAGCATTT GCAGGGGATG ATATGTGTGC TAATAGAAGG 120

TTGCGGCTTA AAACGGAGCA TGAGGGTTTT CTGAACATGA TCTGCCTTAA GGCCAAGGTT 180

CAGTTTGTTT CCCAACCCCAC ATTCTGTGGA TGGTGCTTAT TTAAGGAGGG AATCTTCAAG 240

AAACCTCAAC TAATTTGGGA GCGAATATGC ATAGCCAGAG AGATGGGCAA TCTGGAGAAC 300

TGTATTGACA ATTATGCGAT AGAAGTGTCC TATGCATATA GATTGGGTGA GCTATCAATT 360

GAAATGATGA CAGAAGAAGA AGTGGAGGCA CACTACAATT GTGTGAGGTT CCTGGTTAGG 420

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AACAAGCATA AGATGAGGTG CTCAATT

447

- (2) INFORMATION FOR SEQ ID NO:25:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 149 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:

Glu Ala Ser Thr Phe Leu Phe Asn Thr Met Ala Asn Met Leu Phe Thr 1 5 10 15

Phe Leu Arg Tyr Glu Leu Thr Gly Ser Glu Ser Ile Ala Phe Ala Gly
20 25 30

Asp Asp Met Cys Ala Asn Arg Arg Leu Arg Leu Lys Thr Glu His Glu 35 40 45

Gly Phe Leu Asn Met Ile Cys Leu Lys Ala Lys Val Gln Phe Val Ser 50 55 60

Asn Pro Thr Phe Cys Gly Trp Cys Leu Phe Lys Glu Gly Ile Phe Lys 65 70 75 80

Lys Pro Gln Leu Ile Trp Glu Arg Ile Cys Ile Ala Arg Glu Met Gly 85 90 95

Asn Leu Glu Asn Cys Ile Asp Asn Tyr Ala Ile Glu Val Ser Tyr Ala 100 105 110

Tyr Arg Leu Gly Glu Leu Ser Ile Glu Met Met Thr Glu Glu Glu Val 115 120 125

Glu Ala His Tyr Asn Cys Val Arg Phe Leu Val Arg Asn Lys His Lys 130 135 140

Met Arg Cys Ser Ile 145

- (2) INFORMATION FOR SEQ ID NO:26:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 666 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

ATGAATAATT	TAGTTAAAGC	ATTATCAGCC	TTCGAGTTTA	TAGGTGTTTT	CAATGTGCTC	60
AAATTTCCAG	TTGTTATACA	TAGTGTGCCT	GGTAGTGGTA	AGAGTAGCTT	AATAAGGGAA	120
TTAATCTCAG	AGGACGAGAG	TTTCGTGGCT	TTCACAGCAG	GTGTTCCAGA	CAGTCCTAAC	180
CTCACAGGGA	GGTACATCAA	GCCTTACTCA	CCAGGATGCG	CAGTGCAAGG	AAAAGTGAAT	240
ATACTTGATG	AGTACTTGTC	CGTTCAAGAC	ATTTCGGGTT	TTGATGTACT	GTTTTCAGAC	300
CCGTACCAGA	ATATCAGTAT	TCCCCAAGAG	GCGCATTTCA	TTAAGTCCAA	GACTTGTAGG	360
TTTGGTGTGA	ACACTTGCAA	ATACCTTTCC	TCTTTCGGTT	TCGAAGTTAG	CAGCGACGGG	420
CTGGACGACG	TCATTGTGGG	ATCGCCCTTC	ACTCTAGATG	TTGAAGGGGT	GCTGATATGT	480
TTTGGCAAGG	AGGCGGTAGA	TCTCGCTGTT	GCGCACAACT	CTGAATTCAA	GTTGCCGTGT	540
GAGGTTCGAG	GTTCAACCTT	CAATGTGGTA	ACCCTTTTGA	AATCAAGAGA	CCCAACCCCA	. 600
GAGGACAGGC	ACTGGTTTTA	CATCGCTGCC	ACAAGACATA	GGAAGAAATT	GGTCATTATG	660
CAGTAA						66

(2) INFORMATION FOR SEQ ID NO:27:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 221 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:

Met Asn Asn Leu Val Lys Ala Leu Ser Ala Phe Glu Phe Ile Gly Val 1 5 10 15

Phe Asn Val Leu Lys Phe Pro Val Val Ile His Ser Val Pro Gly Ser 20 25 30

Gly Lys Ser Ser Leu Ile Arg Glu Leu Ile Ser Glu Asp Glu Ser Phe 35 40 45

Val Ala Phe Thr Ala Gly Val Pro Asp Ser Pro Asn Leu Thr Gly Arg 50 55 60

Tyr Ile Lys Pro Tyr Ser Pro Gly Cys Ala Val Gln Gly Lys Val Asn 65 70 75 80

Ile Leu Asp Glu Tyr Leu Ser Val Gln Asp Ile Ser Gly Phe Asp Val 85 90 95

Leu	Phe	Ser	Asp 100	Pro	Tyr	Gln	Asn	Ile 105	Ser	Ile	Pro	Gln	Glu 110	Ala	His
Phe	Ile	Lys 115	Ser	Lys	Thr	Cys	Arg 120	Phe	Gly	Val	Asn	Thr 125	Cys	Lys	Туг
Leu	Ser 130	Ser	Phe	Gly	Phe	Glu 135	Val	Ser	Ser	Asp	Gly 140	Leu	Asp	Asp	Val
Ile 145	Val	Gly	Ser	Pro	Phe 150	Thr	Leu	Asp	Val	Glu 155	Gly	Val	Leu	Ile	Cys 160
Phe	Gly	Lys	Glu	Ala 165	Val	Asp	Leu	Ala	Val 170	Ala	His	Asn	Ser	Glu 175	Phe
Lys	Leu	Pro	Cys 180	Glu	Val	Arg	Gly	Ser 185	Thr	Phe	Asn	Val	Val 190	Thr	Leu
Leu	Lys	Ser 195	Arg	Asp	Pro	Thr	Pro 200	Glu	Asp	Arg	His	Trp 205	Phe	Tyr	Ile
Ala	Ala 210	Thr	Arg	His	Arg	Lys 215	Lys	Leu	Val	Ile	Met 220	Gln			

220

(2) INFORMATION FOR SEQ ID NO:28:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 354 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:28:

ATGCCTTTTC	AGCAGCCTGC	TAATTGGGCA	AAAACCATAA	CTCCATTGAC	TATTGGCTTA	60
GGAATTGGAC	TTGTGCTGCA	TTTTCTGAGA	AAGTCAAATC	TACCATATTC	AGGAGACAAC	120
ATCCATCAAT	TTCCTCACGG	GGGGCGTTAC	CGGGACGGCA	CAAAAAGTAT	AACTTACTGT	180
GGCCCTAAGC	AGTCCTTCCC	CAGTTCAGGA	ATATTTGGTC	AGTCTGAGAA	TTTTGTGCCC	240
TTAATGCTTG	TCATAGGTCT	AATTGCATTC	ATACATGTAT	TGTCTGTTTG	GAATTCTGGT	300
CTTGGTAGGA	ATTGCAATTG	CCATCCAAAT	CCTTGCTCAT	GTAGACAACA	GTAG	354

(2) INFORMATION FOR SEQ ID NO:29:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 116 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi)	SEQU	ENCE	DES	CRIE	OITS	1: SE	Q II	NO:	29:						
Met 1	Pro	Phe	Gln	Gln 5	Pro	Ala	Asn	Trp	Ala 10	Lys	Thr	Ile	Thr	Pro 15	Leu
Thr	Ile	Gly	Leu 20	Gly	Ile	Gly	Leu	Val 25	Leu	His	Phe	Leu	Arg 30	Lys	Ser
Asn	Leu	Pro 35	Tyr	Ser	Gly	Asp	Asn 40	Ile	His	Gln	Phe	Pro 45	His	Gly	СΙΆ
Arg	Tyr 50	Arg	Asp	Gly	Thr	Lys 55	Ile	Thr	Tyr	Cys	Gly 60	Pro	Lys	Gln	Ser
Phe 65	Pro	Ser	Ser	Gly	Ile 70	Phe	Gly	Gln	Ser	Glu 75	Asn	Phe	Val	Pro	Let 80
Met	Leu	Val	Ile	Gly 85	Leu	Ile	Ala	Phe	11e 90	His	Val	Leu	Ser	Val 95	Tr
Asn	Ser	Gly	Leu 100	_	Arg	Asn	Cys	Asn 105	_	His	Pro	Asn	Pro 110	Cys	Se
Cys	Arg	Gln 115													•

(2) INFORMATION FOR SEQ ID NO:30:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 243 base pairs

 - (B) TYPE: nucleic acid (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:

ATGTATTGTC	TGTTTGGAAT	TCTGGTCTTG	GTAGGAATTG	CAATTGCCAT	CCAAATCCTT	60
GCTCATGTAG	ACAACAGTAG	TGGCAGTCAC	CAAGGTTGCT	TTATCAGGGC	CACTGGAGAG	120
TCTATTTTGA	TTGAAAATTG	TGGCCCAAGC	GAGGCCCTTG	CATCAACAGT	GAGGGAGGTG	180
TTGGGGGGTT	TGAAGGCTTT	AGGAATTAGC	CATACTACTG	AAGAAATTGA	TTATCGTTGT	240
TAA						243

(2) INFORMATION FOR SEQ ID NO:31:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 80 amino acids
 - (B) TYPE: amino acid

(C) STRANDEDNESS	:
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(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:31:

Met Tyr Cys Leu Phe Gly Ile Leu Val Leu Val Gly Ile Ala Ile Ala 1 5 10 15

Ile Gln Ile Leu Ala His Val Asp Asn Ser Ser Gly Ser His Gln Gly 20 25 30

Cys Phe Ile Arg Ala Thr Gly Glu Ser Ile Leu Ile Glu Asn Cys Gly
35 40 45

Pro Ser Glu Ala Leu Ala Ser Thr Val Arg Glu Val Leu Gly Gly Leu 50 55 60

Lys Ala Leu Gly Ile Ser His Thr Thr Glu Glu Ile Asp Tyr Arg Cys 70 75 80

(2) INFORMATION FOR SEQ ID NO: 32:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 311 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:

ATGGCGAGTC AAGTTGGTAA GCTCCCCGGA GAATCAAATG AGGCATTTGA AGCCCGGCTG 60

AAATCACTGG AGTTGGCTAG AGCTCAAAAG CAGCCAGAAG GTTCAAACAC ACCGCCTACT 120

CTCAGTGGTG TGCTTGCCAA ACGTAAGAGG GTTATTGAGA ATGCACTCTC AAAGACAGTG 180

GACATGAGGG AGGTGTTGAA ACACGAAACG GTTGTAATTT CCCCAAATGT CATGGATGAG 240

GGTGCAATAG ATGAACTGAT TCGTGCATTC GGAGAATCAG GCATAGCTGA GAGCGCACAA 300

TTTGATGTGG C 311

(2) INFORMATION FOR SEQ ID NO:33:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 103 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:

Met Ala Ser Gln Val Gly Lys Leu Pro Gly Glu Ser Asn Glu Ala Phe 1 5 10 15

Glu Ala Arg Leu Lys Ser Leu Glu Leu Ala Arg Ala Gln Lys Gln Pro 20 25 30

Glu Gly Ser Asn Thr Pro Pro Thr Leu Ser Gly Val Leu Ala Lys Arg
35 40 45

Lys Arg Val Ile Glu Asn Ala Leu Ser Lys Thr Val Asp Met Arg Glu 50 55 60

Val Leu Lys His Glu Thr Val Val Ile Ser Pro Asn Val Met Asp Glu 65 70 75 80

Gly Ala Ile Asp Glu Leu Ile Arg Ala Phe Gly Glu Ser Gly Ile Ala 85 90 95

Glu Ser Ala Gln Phe Asp Val 100

(2) INFORMATION FOR SEQ ID NO:34:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1206 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:

GCAGGATTGA	AGGCTGGCCA	CTGTGTGATT	TTTGATGAGG	TCCAGTTGTT	TCCTCCTGGA	60
TACATCGATC	TATGCTTGCT	TATTATACGT	AGTGATGCTT	TCATTTCACT	TGCCGGTGAT	120
CCATGTCAAA	GCACATATGA	TTCGCAAAAG	GATCGGGCAA	TTTTGGGCGC	TGAGCAGAGT	180
GACATACTTA	GAATGCTTGA	GGGCAAAACG	TATAGGTATA	ACATAGAAAG	CAGGAGGTTT	240
GTGAACCCAA	TGTTCGAATC	AAGACTGCCA	TGTCACTTCA	AAAAGGGTTC	GATGACTGCC	300
GCTTTCGCTG	ATTATGCAAT	CTTCCATAAT	ATGCATGACT	TTCTCCTGGC	GAGGTCAAAA	360
GGTCCTTTGG	ATGCCGTTTT	GGTTTCCAGT	TTTGAGGAGA	AAAAGATAGT	CCAGTCCTAC	420
ヤヤヤにこみないこ	AACAGCTCAC	ACTCACATTI	GGTGAATCAA	CTGGGTTGA	TTTCAAAAAT	480

GGGGGAATTC	TCATATCACA	TGATTCCTTT	CACACAGATG	ATCGGCCGGT	GGCTTACTGC	540
TTTATCTCGC	TTCAGCCACA	ATTTGGATTT	GGTGAACATT	ACAGGTCTGA	GGGTGGAAAG	600
TTTCCTCTCG	CACTTTGCTG	GCAAACCCCT	CTACCATTTT	TTAACAGCCA	AAAGTGGGGA	660
GAATGTCATA	CGAGATTTGC	TCCCAGGTGA	GCCTAACTTC	TTCAGTGGCT	TTAACGTTAG	720
CATTGGAAAG	AATGAAGGTG	TTAGGGAGGA	GAAGTTATGT	GGTGACCCAT	GGTTAAAAGT	780
CATGCTTTTC	CTGGGTCAAG	ATGAGGATTG	TGAAGTTGAA	GAGATGGAGT	CAGAGTGCTC	840
AAATGAAGAA	TGGTTTAAAA	CCCACATTCC	CCTGAGTAAT	CTGGAGTCAA	CCAGGGCTAG	900
GTGGGTGGGT	AAAATGGCTT	TGAAAGAGTA	TCGGGAGGTG	CGTTGTGGTT	ATGAAATGAC	960
TCAACAATTC	TTTGATGAGC	ATAGGGGTGG	AACTGGTGAG	CAACTGAGCA	ATGCATGTGA	1020
GAGGTTTGAA	AGCATTTACC	CAAGGCATAA	AGGAAATGAT	TCAATAACCT	TCCTTATGGC	1080
TGTCCGAAAG	CGTCTCAAAT	TTTCGAAGCC	CCAGGTTGAA	GCTGCCAAAC	TGAGGCGGGC	1140
CAAACCATAT	GGGAAATTCT	TATTAGACTT	TCCTATCCAA	AATCCCATTG	AAAGCCAGTC	1200
ATAATT	-				•	1206

(2) INFORMATION FOR SEQ ID NO:35:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1284 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:

ATTAACCCAA	ATGGTAAGAT	TTCCGCCTTG	TTTGATATAA	CCAATGAGCA	CATAAGGCAT	60
GTTGAGAAGA	TCGGCAATGG	CCCTCAGAGC	ATAAAAGTAG	ATGAGTTGAG	GAAGGTTAAG	120
CGATCCGCCC	TTGATCTTCT	TTCAATGAAT	GGGTCCAAAA	TAACCTATTT	TCCAAACTTT	180
GAGCGGGCTG	AAAAGTTGCA	AGGGTGCTTG	CTAGGGGGCC	TAACTGGTGT	CATAAGTGAT	240
GAAAAGTTCA	GTGATGCAAA	ACCCTGGCTT	TCTGGTATAT	CAACTGCGGA	TATAAAGCCA	300
AGAGAGCTAA	CTGTCGTGCT	TGGCACTTTT	GGGGCTGGAA	AGAGTTTCTT	GTATAAGAGT	360
TTCATGAAGA	GATCTGAGGG	AAAATTTGTA	ACTTTTGTTT	CCCCTAGACG	AGCCTTGGCA	420
AATTCAATCA	AAAATGATCT	TGAAATGGAT	GATGGCTGCA	AAGTTGCCAA	AGCAGGCAAA	480
TCAAAGAAGG	AAGGGTGGGA	TGTAGTGACC	TTTGAAGTTT	TCCTTAGAAA	AGTTTCTGGT	540
TTGAAAGCTG	GTCATTGTGT	GATTTTTGAT	GAGGTTCAGT	TGTTTCCCCC	TGGATACATC	600

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SATCTGTGTT	TACTTGTCAT	ACGAAGTGAT	GCTTTCATTT	CACTTGCTGG	TGATCCATGC	660
CAGAGCACAT	ATGATTCACA	GAAGGATCGA	GCAATTTTGG	GAGCTGAGCA	GAGTGACATA	.720
CTCAGACTGC	TTGAAGGAAA	GACATATAGG	TACAACATAG	AAAGCAGACG	TTTTGTGAAC	780
CCAATGTTTG	AATCTAGACT	ACCATGTCAC	TTCAAAAAGG	GTTCAATGAC	TGCAGCCTTT	840
GCTGATTATG	CAATCTTCCA	CAATATGCAT	GACTTCCTCC	TGGCGAGGTC	AAAAGGCCCC	900
TTGGATGCTG	TTCTAGTTTC	CAGTTTTGAG	GAGAAGAAAA	TAGTCCAATC	CTACTTTGGG	960
ATGAAGCAAC	TCACTCTCAC	ATTTGGTGAA	TCAACTGGGT	TGAACTTCAA	AAATGGAGGA	1020
ATTCTCATAT	CACATGACTC	CTTTCATACT	GACGATCGAC	GGTGGCTTAC	TGCTTTATCT	1080
CGATTCAGCC	ATAATTTGGA	TTTGGTGAAC	ATCACAGGTC	TTGAGGGTGG	AAAGTTTTCT	1140
CTCACATTTT	GCTGGTAAAC	CCCTTTACCA	CTTTTTGACG	GCTTAAAAGT	GGAGAGAATG	1200
TCATACGAGA	CCTGCTTCAG	GTGAGCCTAA	CTTCTTTTAG	GGGTTCAATC	TCAGCATTGG	1260
AAAAAAATGG	AAGGGGTTAG	AGAA				1284

(2) INFORMATION FOR SEQ ID NO:36:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1402 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:

CATTTTTAAA ATTTA	ATCCA GTCGACTCAC	CAAATGTGAG	CGTAAGCTGT	TTCATCCCAA	60
AGTAGGACTG GACTA	ATTTTC TTCTCCTCAP	AACTAGAAAC	CAGAATGGCA	TCCAAAGGAC	120
CTTTTGACCT TGCCA	AGGAGG AAATCATGCA	A TATTGTGGAA	AATGGCATAA	TCAGCAAAGG	180
CAGCAGTCAT TGTAC	CCTTT TTGAAGTGAG	C ATGGCAGTCG	AGATTCAAAC	ATTGGGTTCA	240
CAAATCTTCT GCTTT	CTATG TTGTACCTA	r acgtcttgcc	TTCAAGTATT	TTGAGTATGT	300
CACTCTGCTC AGCGC	CCCAAA ATCGCCCGA	T CTTTTTGTGA	GTCATATGTG	CTCTGACATG	360
GGTCACCAGC AAGTC	GAAATG AAAGCATCA	C TACGTATAAT	AAGCAAACAT	AGATCGATGT	420
ATCCAGGGGG AAACA	AACTGG ACCTCATCG	A AAATTACACA	GTGACCAGCT	TTTAGACCTG	480
CAACTTTTCT AAGG	AAGACT TCAAAAGTC	A CAACATCCCA	TCCTTCCTTC	TTTGACCTGC	540
CTCCTTTGGC AACT	TTGCAG CTATCATCC	A TTTCAAGATO	ATTTTTGATT	GAATTCGCTA	600

GAGCCCGTCT	GGGGGAAACA	AAAGTTACGA	ATTTACCCTC	AGATCTTTTC	ATAAAGCTCT	660
TGTACAAAAA	GCTTTTTCCG	GCTCCAAATG	TGCCAAGCAC	AACAGTTAGC	TCCCTCGGCT	720
TAATGTCAGT	AGTTGATATA	CCAGAAAGCC	AGGGCTTTGC.	ATCACTGAAC	TTCTCATCAC	780
TTATGACACC	AGTTAGGCCT	CCTAGCAGAC	ACCCTTGCAA	CTTTTCAGCC	CGCTCAAAAC	840
TTGGGAAGTA	GGTTACCTTG	GACCCATTAA	TTGAAAGAAG	ATCAAGGGCG	GATCGCTTGA	900
CCTTTCGCAA	TTCATCTACT	TTAATGCTCT	GAGGGCCATT	ACCTATCTTT	TCAACATGCC	960
TTATGTGCTC	ATTAGTTATG	TCAAACAGAG	CGGAAAACTT	GCCATGTGGA	TTAATCACCT	1020
CAATTTCCCC	ATTTATGTCA	CACTTAGCGC	AAATGTCAAA	AGCCTCAAAG	GCTTCAGCTA	1080
AGTTACATCA	TGTTGAGCCT	CCCCCTTGGC	AAAGCTCCTC	AAAAATGTGG	TTAGTGCTAG	1140
GCCTGCACAA	TAATTAACAC	ATCAACTTCA	CCCTGCCAAT	GCTGAACAAT	ACTGTTATCA	1200
TGCAACCATC	CATGGGGCAC	ATGGTTGGAA	TTGATTGATT	TAAGGCAAAA	ATCCCCACAG	1260
GGGGCATCCC	CTTCCCCAAT	TTCCACTGAT	TCATACTCTG	GCGTTATCAT	ATCAACCCAA	1320
TGTGTCAAAT	ACAAATAATG	CAATCTCTCA	TCTCCGATAA	CATTTCCCCC	ATTTTTTAAA	1380
AATGGTGGGG	TGAAAATTGG	AA	•			1402

(2) INFORMATION FOR SEQ ID NO:37:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1236 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

GTGGTTTTTG	CAACAACAGG	CCCAGGTCTA	TCTAAGGTTT	TGGAAATGCC	TCGAAGCAAG	60
AAGCAATCTA	TTCTGGTTCT	TGAGGGAGCC	CTATCCATAG	AAACGGACTA	TGGCCCAAAA	120
GTTCTGGGAT	CTTTTGAAGT	TTTCAAAGGG	GATTTCAACA	TTAAAAAAATT	GGAAGAAGT	180
TCCATCTTTG	TAATAACATA	CAAGGCCCCA	GTTAGATCTA	CTGGCAAGTT	GAGGGTCCAC	240
CAATCAGAAT	GCTCATTTTC	TGGATCCAAG	GAGGTATTGC	TGGGTTGTCA	GATTGAGGCA	300
TGTGCTGATT	ATGATATTGA	TGATTTCAAT	ACTTTCTTTG	TACCTGGTGA	TGGTAATTGC	360
TTTTGGCATT	CAGTTGGTTT	CTTACTCAGT	ACTGACGGAC	TTGCTTTGAA	GGCCGGCATT	420
CGTTCTTTCG	TGGAGAGTGA	ACGCCTGGTG	AGTCCAGATC	TTTCAGCCCC	AACCATTTCT	480
AAACAACTGG	GGGAAAATGC	TTATGCCGAG	AATGAGATGA	TTGCATTATT	TTGTATTCGA	540

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GGCTGATAGT	GATTACGCCA	GAGTATGAAG	TCAGTTGGAA	ATTTGGGGAA	600
CCCTGTGCGG	AATTCTTTGC	CTTAAATCAA	ATCACTTCCA	ACCATGTGCC	660
GTTGCATGAT	TACAGCTATT	GCTTCAGCAC	TTGGTAGGCG	TGAAGTTGAT	720
ATCTGTGCAG	GCCTAGCACT	AACCACATTT	TTGAGGAGCT	TTGCCAAGGG	780
ACATGATGTA	CTTAGCTGAA	GCCTTTGAGG	CTTTTGACAT	TTGCGCTAAG	840
ATGGGGAAAT	TGAGGTGATT	AATCCACATG	GCAAGTTTTC	CGCTCTGTTT	900
ATGAGCACAT	AAGGCATGTT	GAAAAGATAG	GTAATGGCCC	TCAGAGCATT	960
AATTGCGAAA	GGTCAAGCGA	TCTGCCCTTG	ATCTTCTTTC	AATTAATGGG	1020
CCTACTTCCC	AAGTTTTGAG	CGGGCTGAAA	AGTTGCAAGG	GTGTCTGCTA	1080
CTGGTGTCAT	AAGTGATGAG	AAAGTCAGTG	ATGCAAAGC	CTGCTTTTTG	1140
TACTGACATT	AAGCCGAGGG	AGCTAACTGT	TGTGCTTTG	G CACATTTGGA	120
AGCCTTTTG1	ACCAAGAGCT	TTATTG			123
	CCCTGTGCGG GTTGCATGAT ATCTGTGCAG ACATGATGTA ATGGGGAAAT ATGAGCACAT AATTGCGAAA CCTACTTCCC CTGGTGTCAT TACTGACATT	CCCTGTGCGG AATTCTTTGC GTTGCATGAT TACAGCTATT ATCTGTGCAG GCCTAGCACT ACATGATGTA CTTAGCTGAA ATGGGGAAAT TGAGGTGATT ATGAGCACAT AAGGCATGTT AATTGCGAAA GGTCAAGCGA CCTACTTCCC AAGTTTTGAG CTGGTGTCAT AAGTGATGAG TACTGACATT AAGCCGAGGG	CCCTGTGCGG AATTCTTTGC CTTAAATCAA GTTGCATGAT TACAGCTATT GCTTCAGCAC ATCTGTGCAG GCCTAGCACT AACCACATTT ACATGATGTA CTTAGCTGAA GCCTTTGAGG ATGGGGAAAT TGAGGTGATT AATCCACATG ATGAGCACAT AAGGCATGTT GAAAAGATAG AATTGCGAAA GGTCAAGCGA TCTGCCCTTG CCTACTTCCC AAGTTTTGAG CGGGCTGAAA CCTGGTGTCAT AAGTGATGAG AAAGTCAGTG	CCCTGTGCGG AATTCTTTGC CTTAAATCAA ATCACTTCCA GTTGCATGAT TACAGCTATT GCTTCAGCAC TTGGTAGGCG ATCTGTGCAG GCCTAGCACT AACCACATTT TTGAGGAGCT ACATGATGTA CTTAGCTGAA GCCTTTGAGG CTTTTGACAT ATGGGGAAAT TGAGGTGATT AATCCACATG GCAAGTTTTC ATGAGCACAT AAGGCATGTT GAAAAGATAG GTAATGGCCC AATTGCGAAA GGTCAAGCGA TCTGCCCTTG ATCTTCTTTC CCTACTTCCC AAGTTTTGAG CGGGCTGAAA AGTTGCAAGGG CTGGTGTCAT AAGTGATGAG AAAGTCAGTG ATGCAAAGCCC TACTGACATT AAGCCGAGGG AGCTAACTGT TGTGCTTTTGCCCTTGCCCTTG ATCTTCTTTCCCCCTTGCCCTTG ATCTTCTTTCCCCCTTGCCCTTGCCCTTGCCCTTGCCCAAGGCCCCCCCC	GGCTGATAGT GATTACGCCA GAGTATGAAG TCAGTTGGAA ATTTGGGGAA CCCTGTGCGG AATTCTTTGC CTTAAATCAA ATCACTTCCA ACCATGTGCC GTTGCATGAT TACAGCTATT GCTTCAGCAC TTGGTAGGCG TGAAGTTGAT ATCTGTGCAG GCCTAGCACT AACCACATTT TTGAGGAGCT TTGCCAAGGG ACATGATGTA CTTAGCTGAA GCCTTTGAGG CTTTTGACAT TTGCGCTAAG ATGGGGAAAT TGAGGTGATT AATCCACATG GCAAGTTTTC CGCTCTGTTT ATGAGCACAT AAGGCATGTT GAAAAGATAG GTAATGGCCC TCAGAGCATT AATTGCGAAA GGTCAAGCGA TCTGCCCTTG ATCTTCTTC AATTAATGGG CCTACTTCCC AAGTTTTGAG CGGGCTGAAA AGTTGCAAGG GTGTCTGCTA CTGGTGTCAT AAGTGATGAG AAAGTCAGTG ATGCAAAGCC CTGCTTTTTG TACTGACATT AAGCCGAGGG AGCTAACTGT TGTGCTTTGG CACATTTGGA AGCCTTTTGT ACCAAGAGCT TTATTG

(2) INFORMATION FOR SEQ ID NO:38:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1296 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:38:

GTCTAACTGG CGTTATAAGT GATGAGAAAT TCAGTGATGC AAAACCTTGG CTTTCTGGTA 60 TATCTACTAC AGATATTAAG CCAAGGGAAT TAACTGTTGT GCTTGGTACA TTTGGGGCTG 120 GGAAGAGTTT CTTGTACAAG AGTTTCATGA AAAGGTCTGA GGGTAAATTC GTAACCTTTG 180 TTTCTCCCAG ACGTGCTTTA GCAAATTCAA TCAAAAATGA TCTTGAAATG GATGATAGCT 240 GCAAAGTTGC CAAAGCAGGT AGGTCAAAGA AGGAAGGGTG GGATGTAGTA ACTTTTGAGG 300 TCTTCCTCAG AAAAGTTGCA GGATTGAAGG CTGGCCACTG TGTGATTTTT GATGAGGTCC 360 AGTTGTTTCC TCCTGGATAC ATCGATCTAT GCTTGCTTAT TATACGTAGT GATGCTTTCA 420 TTTCACTTGC CGGTGATCCA TGTCAAAGCA CATATGATTC GCAAAAGGAT CGGGCAATTT 480 TGGGCGCTGA GCAGAGTGAC ATACTTAGAA TGCTTGAGGG CAAAACGTAT AGGTATAACA 540 TAGAAAGCAG GAGGTTTGTG AACCCAATGT TCGAATCAAG ACTGCCATGT CACTTCAAAA 600

AGGGTTCGAT	GACTGCCGCT	TTCGCTGATT	ATGCAATCTT	CCATAATATG	CATGACTTTC	660
TCCTGGCGAG	GTCAAAAGGT	CCTTTGGATG	CCGTTTTGGŢ	TTCCAGTTTT	GAGGAGAAAA	720
AGATAGTCCA	GTCCTACTTT	GGAATGAAAC	AGCTCACACT	CACATTTGGT	GAATCAACTG	780
GGTTGAATTT	CAAAAATGGG	GGAATTCTCA	TATCACATGA	TTCCTTTCAC	ACAGATGATC	840
GGCGGTGGCT	TACTGCTTTA	TCTCGCTTCA	GCCACAATTT	GGATTTGGTG	AACATTACAG	900
GTCTGAGGTG	GAAAGTTTCC	TCTCGCACTT	TGCTGGCAAA	CCCCTCTACC	ATTTTTTAAC	960
AGCCAAAAGT	GGGGAGAATG	TCATACGAGA	TTTGCTCCCA	GGTGAGCCTA	ACTTCTTCAG	1020
TGGCTTTAAC	GTTAGCATTG	GAAAGAATGA	AGGTGTTAGG	GAGGAGAAGT	TATGTGGTGA	1080
CCCATGGTTA	AAAGTCATGC	TTTTCCTGGG	TCAAGATGAG	GATTGTGAAG	TTGAAGAGAT	1140
GGAGTCAGAG	TGCTCAAATG	AAGAATGGTT	TAAAACCCAC	ATTCCCCTGA	GTAATCTGGA	1200
GTCAACCAGG	GCTAGGTGGG	TGGGTAAAAT	GGCCTTGAAA	GAGTATCGGG	AGGTGCGTTG	1260
TGGTTATGAA	ATGACTCAAC	AATTCTTTGA	TGACAT			1296

(2) INFORMATION FOR SEQ ID NO:39:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 907 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:39:

ATGTTCACCA	AATCCAAATT	ATGGCTGAAG	CGAGATAAAG	CAGTAAGCCA	CCGCCGATCA	60
TCTGTGTGA	AGGAATCATG	TGATATGAGA	ATTCCCCCAT	TTTTGAAATT	CAACCCAGTT	120
GATTCACCA	ATGTGAGTGT	GAGCTGTTTC	ATTCCAAAGT	AGGACTGGAC	TATCTTTTTC	180
TCCTCAAAA	TGGAAACCAA	AACGGCATCC	AAAGGACCTT	TTGACCTCGC	CAGGAGAAAG	240
TCATGCATAT	TATGGAAGAT	TGCATAATCA	GCGAAAGCGG	CAGTCATTGA	GCCCTTTTTG	300
AATTGACATO	GCAGTCTTGA	TTCGAACATT	GGATTCACAA	ACCTCCTGCT	TTCAATGTTA	360
TACCTATACO	TCTTGCCCTC	AAGCAGTCTA	AGTATGTCAC	TCTGCTCAGC	GCCCAAAATT	420
GCCCGATCCT	TTTGCGAATC	ATATGTGCTT	TGACATGGAT	CACCGGCAAG	TGAAATGAAA	480
GCATCACTAC	CGTATAATAAG	CAAGCATAGA	TCGATGTATC	CAGGAGGAAA	CAACTGGACC	540
TCATCGAAA	A TCACACAGTG	GCCAGCCTTC	AATCCTGCAA	CTTTTCTGAG	GAAAACCTCA	600
AAAGTTACT	CATCCCACCC	TTCCTTCTTT	GACCTACCTG	CTTTAGCAAC	TTTGCAGCTA	660

	CAAGATCATT		+ .	CTTCCCAGCC	720
•				AGATATACCA	840
	•			TAGGCCCCCT	900
AGCAAAC	GIIIIGCAIC	ACTOMICTIC		,	90
AGCMMAC					

(2) INFORMATION FOR SEQ ID NO:40:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1232 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:

AGAATGCTTA TGCTGAGAAT GAGATGATTG CATTATTTTG CATCCGGCAC CATGTAAGGC 60 TTATAGTAAT AACACCGGAA TATGAAGTTA GTTGGAAATT TGGGGAAAGT GAGTGGCCCC 120 TATGTGGAAT TCTTTGCCTG AGGTCCAATC ACTTCCAACC ATGCGCCCCG CTGAATGGTT 180 GCATGATCAC GGCTATTGCT TCAGCACTTG GGAGGCGTGA GGTTGATGTG TTAAATTATC 240 TGTGTAGGCC TAGCACTAAT CACATCTTTG AGGAGCTGTG CCAGGGCGGA GGGCTTAATA 300 TGATGTACTT GGCTGAAGCT TTTGAGGCCT TTGACATTTG TGCAAAGTGC GACATAAATG 360 GGGAAATTGA GGTCATTAAC CCAAATGGCA AGATTTCCGC CTTGTTTGAT ATAACTAATG 420 AGCACATAAG GCATGTTGAG AAGATCAGCA ATGGCCCTCA GAGCATAAAA ATAGATGAGT 480 TGAGGAAGGT TAAGCGATCC CGCCTTGACC TTCTTTCAAT GAATGGGTCC AAAATAACCT 540 ATTTTCCAAA CTTTGAGCGG GCTGAAAAGT TGCAAGGGTG CTTGCTAGAG GGCCTGACTG 600 GTGTCATAAG TGATGAAAAG TTCAGTGATG CAAAACCTTG GCTTTCTGGT ATATCAACTG 660 720 CGGATATTAA GCCAAGAGAG CTAACTGTCG TGCTTGGCAC ATTTGGTGCT GGAAAGAGTT TCTTGTATAA GAGTTTCATG AAGAGATCTG AAGGAAAATT TGTAACTTTT GTTTCCCCTA 780 840 GGCGAGCTTT GGCCAATTCG ATCAAGAATG ATCTTGAAAT GGATGATGGC TGCAAAGTTG CCAAAGCAGG CAAGTCAAAG AAGGAAGGGT GGGATGTGGT AACATTTGAG GTTTTCCTTA 900 GAAAAGTTTC TGGTTTGAAG GCTGGTCATT GTGTGATTTT CGATGAGGTT CAGTTGTTTC 960 CCCCTGGATA TATCGATCTA TGTTTACTTG TCATACGCAG TGATGCTTTT ATTTCACTTG 1020 WO 98/52964

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CCGGTGA	ATCC ATGCCAGAGC ACATATGATT CACAAAAGGA TCGGGCAA	TT TTGGGAGCTG	1080
AGCAGAG	STGA CATACTCAGA TTGCTTGAAG GAAAGACGTA TAGGTACAA	AC ATAGAAAGCA	1140
GACGTŤT	TTGT GAACCCAATG TTTGAATTTA GACTACCATG TCACTTCA	AA AAAGGGTTCA	1200
ATGACTO	GCTG CCTTTGCTGA TTATGCAATC TT		1232
(2) INF	FORMATION FOR SEQ ID NO:41:		
(i	i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear		
(ii	i) MOLECULE TYPE: cDNA		
(xi	i) SEQUENCE DESCRIPTION: SEQ ID NO:41:	,	,
GCTTCAG	GCAC TTGGAAGGCG		20
(2) INF	FORMATION FOR SEQ ID NO:42:		
(i	i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear		
(ii	i) MOLECULE TYPE: cDNA		
(x)	i) SEQUENCE DESCRIPTION: SEQ ID NO:42:		
CACACAG	GTGG CCAGCCT		17
(2) IN	FORMATION FOR SEQ ID NO:43:		
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear		
(i:	i) MOLECULE TYPE: cDNA		

(2) INFORMATION FOR SEQ ID NO:44:

GGAGGTGCGT TGTGGTTATG

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
		,
ix)	i) SEQUENCE DESCRIPTION: SEQ ID NO:44:	
CCCTGGG	CACT GCACACCC	18
(2) IN	FORMATION FOR SEQ ID NO:45:	
(:	i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(i	i) MOLECULE TYPE: cDNA	
(x	i) SEQUENCE DESCRIPTION: SEQ ID NO:45:	
GGAGGT	GACC ACATTACG	18
(2) IN	FORMATION FOR SEQ ID NO:46:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear ii) MOLECULE TYPE: cDNA	
		•
. (:	xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:	
CATCA	CGACT TGTCACAAAC C	2
(2) I	NFORMATION FOR SEQ ID NO:47:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	

GGCCAAGGTT CAGTTTG

(2) INFORMATION FOR SEQ ID NO:51:

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(xi) SEQUENCE DESCRIPTION: SEQ II	NO:47:
TGGGCCTCCA CTTCTTC	17
(2) INFORMATION FOR SEQ ID NO:48:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ I	D NO:48:
GGGGTTGCCT GAAGAT	16
(2) INFORMATION FOR SEQ ID NO:49:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ I	D NO:49:
ACACCTGCTG TGAAAGC	17
(2) INFORMATION FOR SEQ ID NO:50:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: cDNA	
(xi) SEQUENCE DESCRIPTION: SEQ	ID NO:50:

(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 20 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear			
(ii) MOLECULE TYPE: cDNA			
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:51: GATGAGGTCC AGTTGTTTCC (2) INFORMATION FOR SEQ ID NO:52:			20
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear			
(ii) MOLECULE TYPE: cDNA			
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:			20
ATCCAAAGGA CCTTTTGACC			_
(2) INFORMATION FOR SEQ ID NO:53:		·	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear			
(ii) MOLECULE TYPE: cDNA			
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:	•		17
CTTGATGAGT ACTTGTC			1.
(2) INFORMATION FOR SEQ ID NO:54:			
(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 17 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear			
(ii) MOLECULE TYPE: cDNA			

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:

GCAAGGATTT GGATGGC

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WHAT IS CLAIMED:

- 1. An isolated protein or polypeptide corresponding to a protein or polypeptide of a Rupestris stem pitting associated virus.
- 2. The isolated protein or polypeptide according to claim 1, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 3. The isolated protein or polypeptide according to claim 2, wherein the protein or polypeptide is a replicase.
- 4. The isolated protein or polypeptide according to claim 3, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 3, SEQ. ID. No. 14, or SEQ. ID. No. 25.
- 5. The isolated protein or polypeptide according to claim 3, wherein the protein or polypeptide has a molecular weight of about 240 to 246 kDa.
- 6. The isolated protein or polypeptide according to claim 2, wherein the protein or polypeptide is a coat protein.
- 7. The isolated protein or polypeptide according to claim 6, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 11, SEQ. ID. No. 22, or SEQ. ID. No. 33.
- 8. The isolated protein or polypeptide according to claim 6, wherein the protein or polypeptide has a molecular weight of about 25 to 30 kDa.
- 9. The isolated protein or polypeptide of claim 2, wherein the protein or polypeptide is a protein of a triple gene block.

- 10. The isolated protein or polypeptide according to claim 9, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 5, SEQ. ID. No. 16, or SEQ. ID. No. 27.
- 11. The isolated protein or polypeptide according to claim 9, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 7, SEQ. ID. No. 18, or SEQ. ID. No. 29.
- 12. The isolated protein or polypeptide according to claim 9, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 9, SEQ. ID. No. 20, or SEQ. ID. No. 31.
- 13. The isolated protein or polypeptide according to claim 9, wherein the protein or polypeptide has a molecular weight of 20 to 26 kDa, 10 to 15 kDa, or 5 to 10 kDa.
- 14. The isolated protein or polypeptide according to claim 1, wherein the protein or polypeptide is purified.
- 15. The isolated protein or polypeptide according to claim 1, wherein the protein or polypeptide is recombinant.
- 16. An isolated RNA molecule encoding a protein or polypeptide according to claim 1.
- 17. The isolated RNA molecule according to claim 16, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 18. An isolated DNA molecule encoding a protein or polypeptide according to claim 1.

- 19. The isolated DNA molecule according to claim 18, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 20. The isolated DNA molecule according to claim 19, wherein the protein or polypeptide is a replicase.
- 21. The isolated DNA molecule according to claim 20, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 3, SEQ. ID. No. 14, or SEQ. ID. No. 25.
- 22. The isolated DNA molecule according to claim 21, wherein the DNA molecule comprises a nucleotide sequence corresponding to SEQ. ID. No. 2, SEQ. ID. No. 13, or SEQ. ID. No. 24.
- 23. The isolated DNA molecule according to claim 19, wherein the protein or polypeptide is a coat protein.
- 24. The isolated DNA molecule according to claim 23, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 11, SEQ. ID. No. 22, or SEQ. ID. No. 33.
- 25. The isolated DNA molecule according to claim 24, wherein the DNA molecule comprises a nucleotide sequence corresponding to SEQ. ID. No. 10, SEQ. ID. No. 21, or SEQ. ID. No. 32.
- 26. The isolated DNA molecule according to claim 19, wherein the protein or polypeptide is a protein of a triple gene block.
- 27. The isolated DNA molecule according to claim 26, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 5, SEQ. ID. No. 16, or SEQ. ID. No. 27.

- 28. The isolated DNA molecule according to claim 27, wherein the DNA molecule comprises a nucleotide sequence corresponding to SEQ. ID. No. 4, SEQ. ID. No. 15, or SEQ. ID. No. 26.
- 29. The isolated DNA molecule according to claim 26, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 7, SEQ. ID. No. 18, or SEQ. ID. No. 29.
- 30. The isolated DNA molecule according to claim 29, wherein the DNA molecule comprises a nucleotide sequence corresponding to SEQ. ID. No. 6, SEQ. ID. No. 17, or SEQ. ID. No. 28.
- 31. The isolated DNA molecule according to claim 26, wherein the protein or polypeptide comprises an amino acid sequence corresponding to SEQ. ID. No. 9, SEQ. ID. No. 20, or SEQ. ID. No. 31.
- 32. The isolated DNA molecule according to claim 31, wherein the DNA molecule comprises a nucleotide sequence corresponding to SEQ. ID. No. 8, SEQ. ID. No. 19, or SEQ. ID. No. 30.
- 33. An expression system comprising a vector into which is incorporated a heterologous DNA molecule according to claim 18.
- 34. The expression system according to claim 33, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 35. A host cell transformed with a heterologous DNA molecule according to claim 18.
- 36. The host cell according to claim 35, wherein the host cell is selected from a group consisting of Agrobacterium vitis and Agrobacterium tumefaciens.

- 37. The host cell according to claim 35, wherein the host cell is a grape cell.
- 38. The host cell according to claim 35, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 39. A transgenic *Vitis* scion cultivar or rootstock cultivar comprising the DNA molecule according to claim 18.
- 40. A transgenic *Vitis* scion cultivar or rootstock cultivar according to claim 39, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 41. A method of imparting *Rupestris* stem pitting associated virus resistance to a *Vitis* scion cultivar or rootstock cultivar comprising:

transforming a *Vitis* scion cultivar or rootstock cultivar with a DNA molecule according to claim 18.

- 42. The method according to claim 41, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 43. The method according to claim 41, wherein the *Rupestris* stem pitting associated virus is RSPaV-1, RSP47-4, or RSP158.
- 44. The method according to claim 41, wherein said transforming is *Agrobacterium* mediated.

45. The method according to claim 41, wherein said transforming comprises:

propelling particles at grape plant cells under conditions effective for the particles to penetrate into the cell interior and

introducing an expression vector comprising the DNA molecule into the cell interior.

- 46. An antibody or binding portion thereof or probe recognizing the protein or polypeptide according to claim 1.
- 47. The antibody or binding portion thereof or probe according to claim 46, wherein the protein or polypeptide is selected from a group consisting of a replicase, a coat protein, and a protein of a triple gene block.
- 48. A method for detection of *Rupestris* stem pitting associated virus in a sample, said method comprising:

providing an antibody or binding portion thereof recognizing the protein or polypeptide according to claim 1;

contacting the sample with the antibody or binding portion thereof; and detecting any reaction which indicates that *Rupestris* stem pitting associated virus is present in the sample using an assay system.

- 49. A method according to claim 48, wherein the assay system is selected from a group consisting of enzyme-linked immunoabsorbent assay, radioimmunoassay, gel diffusion precipitin reaction assay, immunodiffusion assay, agglutination assay, fluorescent immunoassay, protein A immunoassay, and immunoelectrophoresis assay.
- 50. A method according to claim 48, wherein said detecting is effective to detect any strain of *Rupestris* stem pitting associated virus.

51. A method for detection of *Rupestris* stem pitting associated virus in a sample, said method comprising:

providing a nucleotide sequence of the DNA molecule according to claim 18 as a probe in a nucleic acid hybridization assay;

contacting the sample with the probe; and
detecting any reaction which indicates that *Rupestris* stem pitting associated virus is present in the sample.

- 52. A method according to claim 51, wherein the nucleic acid hybridization assay is selected from a group consisting of dot blot hybridization, tissue printing, southern hybridization, and northern hybridization.
- 53. A method according to claim 51, wherein said detecting is effective to detect any strain of *Rupestris* stem pitting associated virus.
- 54. A method according to claim 53, wherein the probe has a nucleotide sequence selected from a group consisting of SEQ. ID. No. 53, SEQ. ID. No. 54, SEQ. ID. No. 51, and SEQ. ID. No. 52.
- 55. A method for detection of *Rupestris* stem pitting associated virus in a sample, said method comprising:

providing a nucleotide sequence of the DNA molecule according to claim 18 as a probe in a gene amplification detection procedure;

contacting the sample with the probe; and
detecting any reaction which indicates that *Rupestris* stem pitting associated virus is present in the sample.

- 56. A method according to claim 55, wherein the gene amplification detection procedure is selected from a group consisting of polymerase chain reaction and immunocapture polymerase chain reaction.
- 57. A method according to claim 55, wherein said detecting is effective to detect any strain of *Rupestris* stem pitting associated virus.

- 58. A method according to claim 57, wherein the probe has a nucleotide sequence selected from a group consisting of SEQ. ID. No. 53, SEQ. ID. No. 54, SEQ. ID. No. 51, and SEQ. ID. No. 52.
- 59. An oligonucleotide primer capable of hybridizing to a nucleic acid of a *Rupestris* stem pitting associated virus.
- 60. An oligonucleotide primer according to claim 59, wherein the oligonucleotide primer comprises a nucleotide sequence of SEQ. ID. No. 41, SEQ. ID. No. 42, SEQ. ID. No. 43, SEQ. ID. No. 44, SEQ. ID. No. 45, SEQ. ID. No. 46, SEQ. ID. No. 47, SEQ. ID. No. 48, SEQ. ID. No. 49, SEQ. ID. No. 50, SEQ. ID. No. 51, SEQ. ID. No. 52, SEQ. ID. No. 53, or SEQ. ID. No. 54.
- 61. An oligonucleotide primer according to claim 59, wherein the oligonucleotide primer is capable of hybridizing to a nucleic acid of any strain of *Rupestris* stem pitting associated virus and comprises a nucleotide sequence of SEQ. ID. No. 51, SEQ. ID. No. 52, SEQ. ID. No. 53, or SEQ. ID. No. 54.
- 62. The isolated DNA molecule according to claim 18, wherein the DNA molecule comprises a nucleotide sequence of SEQ. ID. No. 34, SEQ. ID. No. 35, SEQ. ID. No. 36, SEQ. ID. No. 37, SEQ. ID. No. 38, SEQ. ID. No. 39, SEQ. ID. No. 40.
- 63. The isolated DNA molecule according to claim 18 wherein the DNA molecule comprises a nucleotide sequence of SEQ. ID. No. 1, SEQ. ID. No. 12, or SEQ. ID. No. 23.



FIG. 1

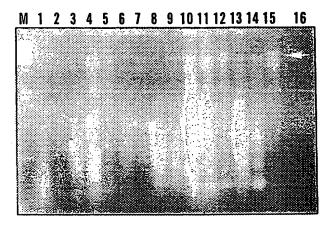


FIG. 2A

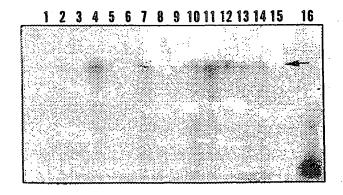
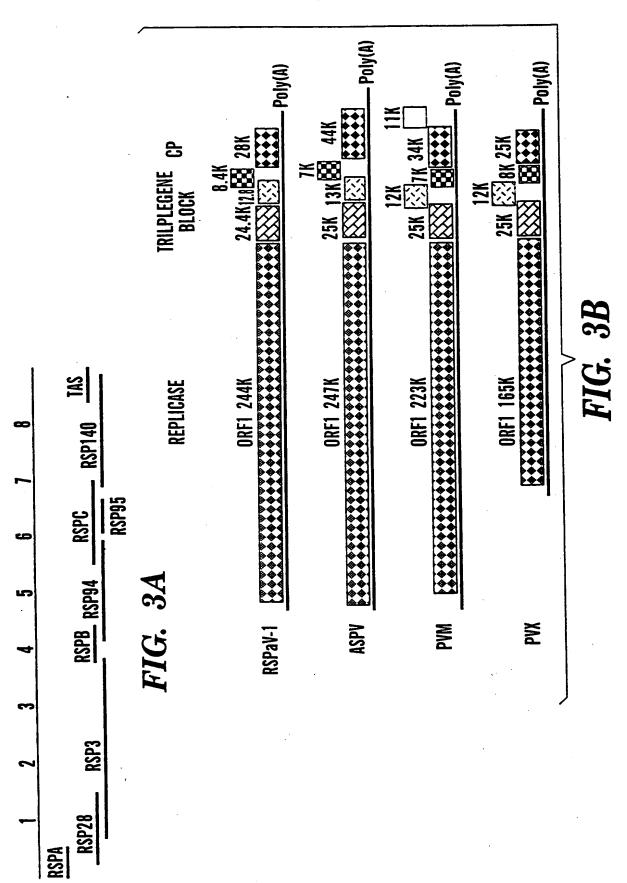


FIG. 2B

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MA+. #R. +. E + +. +. F + . +. + Q + +. +	HS HP VCKTLEN. I L# N# LPS Y # * S F Y# V# I K K# * LK * . # * L V * NR. # # S. D RY * F # HS HP VCKTLENYI LYS VLPS YI - NS S F YF VGI KERKLQLLKS KCKNLDS VQVVNRYVTS ADRMRYTNDF V HS HP VCKTLENNI LF NI LPS YL - DNS F YL VSI KKNKVDFLKRRHPDLQMVETI NRYI SSI DKTRYGGF F HHS HP VCKTLENHI LYNVLPS YV - NNS F YF VGI KDFKLQFLKRRNKDLS LVALI NRF VTS RDVS RYGS EF V	* S *	V. PPE. L. G. ESLNPW. Y. Y. I. G. L. F. PDG E. Y. QPL YLL. ARS LPDG Y. VD vvyp pellfkptrslnewcytydi vgdtlmffpdgvqsegyqqplkggyllgarslklpdgtvymvdvlc i vfppei lagakeslnpwcytfri vgkdlvffpdgeqseayi Qpvagsyllrtgki ttpsgdi fqldllkavi ppevlvgspeslnpwayqyki ngnqllfapdgnwnemysqplscryllkarsvvlpdgsrysvdi i h	S#F*HHL.S.TG*R.F.*F.A*.L.*.L#**#P###.KIY.YLRTLKKPD SKFPHHLISIT-KGEAAAPTHRAFGPFEAVASEALKATLSPDYPCAFPVSYEVVNKIYRYLRTLKKPDEQ SSFSHHLISIT-KGEAIGQKMRFFNGFEAVAMKGLNP-LRRKVESCLPISKNTILKIYRYLRTLKKPDLQ SSFSHHLISIT-KGEAIGQKMRFFNGFEAVAMKGLNP-LRRKVESCLPISKNTILKIYRYLRTLKKPDLQ SKFSHHLLSFTPMGNLLTSNMRCFSGFDAIGIKDLEP-LSRGMHSCFPVHHDVVTKIYLYLRTLKKPDKE	SA. AKL. Q. # P. G#EI * F. E. F *. L# # SAI AKLSQI I AEPSGREI DFVECFARLVI (371) SAMAKLSQVCKDPNGYEI KFFEEFSKLCL (373) SAMAKLSQVCKDPNGYEI KFFEEFSLVI (372)
Consensus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1	Consersus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1	Consensus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1	Consersus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1	Consersus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1	Consersus PVM Rep-1 ASPV Rep-1 RSPaV-1 Rep-1

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+GTFG. GKS. L K. +. +. GK + FVSPRR#LA. • + #. • • . # # K#G. • • V. T • E. (1163)1 VGTFGSGKSTLF- KNLLKYGAGKSLDFVSPRRALAEDFKRTVGMNERGGRAKAGGE NWRVTTLET (1372)1 LGTFGCGKSSLF- KKFI EKSPGKAI TFVSPRRSLAESI NHDLGLARVGGK- KTGKSKDLKNVRVKTFEL (1354)VLGTFGAGKSFLY- KSFMKRSEGKFVTFVSPRRALANSI KNDLEMDDSCKVAKAGRSKKEGWD- VVTFEV	F## FLARV FILHL FLRKV	L. * Y. # * *. S. RF. N*. F RLPC K T * # * # # * VLVSSF*EK* LGAREYNYKVRSHRFLNCNFIGRLPCEINKDDCTIDEPHIMRMHL-BNLLDVAEEYKSVVLVSSFDEKM LSGRSYKFNILSQRFRNPVFYGRLPCNLNKTRLTLDEEEYTLWDSIQEFSMMGRKDCPV-VLVSSFEEKK -II LEGKTYRYNIESRRFVNPMFESRLPCHFKKGSMTAAFADYAIFHNMHDFLLARSKGPLDAVLVSSFEEKK	*V**T#GESTGL*FG.I*##ST.*RRW.TAL*RF**###NG** vvcahlpea-kvltfgestgltfmhgti yi savsertnerrwi talrrfrhucfvncsgmdyqqlagry i vaahlglkmkci tygestglnfqkgai Lytyesaltsdrrwwtalsrfshdi hfi ngmgvtwdnai thf i vqsyfgmkqltltfgestglnfknggi Li shdsfhtddrrwtalsrfshnldlvni tglrvesflshf	. G***F#*	PEVVMQEEWFRTHLPRDELESVRAQWVHKILAKEYREWRMGDMVSEQFTHDI PEVVMQEEWFRTHLPRDELESVRAQWVHKLISREDREFRIGDITTEQFTHDI DEVEAAEDWFKTHIPIMSLEAVRAQWVHKLISREDREFRIGDITTEQFTDDI ESECSNEEWFKTHIPLSNLESTRARWVGKMALKEYREVRCGYEMTQQFFDEI
Ccrisersus PVM Rep-11 ASPV Rep-11 RSPaV-1 Rep-11	Corsensus PVM Rep-11 ASPV Rep-11 RSPaV-1 Rep-11	Conseins PVM Rep-11 ASPV Rep-11 RSPaV-1 Rep-11	Consensus PVM Rep-II ASPV Rep-II RSPaV-I Rep-II	Conscisus PVM Rep-11 ASPV Rep-11 RSPaV-1 Rep-11	Corsersus PVM Rep-II ASPV Rep-II RSPaV-I Rep-II

WNF HAF

YPRH**. D. . TFLMAV. KRL. FS. P. . E. **L**A**#GK*LL*. FL. . . PL#. *H#.. *. . EA*-. YPRHRAS DTYTFLMAVKRRLS FS NPGKEKGNLF HAAS YGKALLS EFLKR VPLKP NHNVR F MEEAL- YPRHKGTDTATFLMAVKRRLS FS SPAAEHAKLRRAKPFGKFLLDTFLKR VPLNS SHDEKMMQEAV- YPRHKGNDSI TFLMAVKRRLKFS KPQVEAAKLRRAKPYGKFLLDS FLS KIPLKAS HNSI MFHEAV.

RSPaV-1 Rep-II

ASPV Rep-1

PVM Rep-II

CONSCINSUS

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E. KK. S KS # ATI ENH. GRS C# DW. . D# A*I F * KS Q. CTKF DNR- . R# AKA* Q** * CF QH# VL. RF AP YMR EEKKLS KS AATI ENHS GRS CR DWP T DV AQI F S KS QL CTKF DNR- FRVAKAAQSI VCF QHA VL CRF AP YMR EEKKLS KS MATI ENHS GRS CE DWP V DKALI F MKS QL CTKF DNR- FRS AKAGQTL ACF QHS VL CRF AP YMR EAKKAS KS AATI ENHAGRS CR DWLL DVALI F MKS QHCTKF DNR- LR VAKAGQTL ACF QHA VL VRF AP YMR RSPaV-1 Rep-II ASPV Rep-11 PVM Rep-II Consensus

. K...L.N. YI HSGK##j*#L..WV. *...F. *#CTESDYEAFDASQD*FI *AFEL..

YI EMKVHEVLPKNYYI HSGKGLEELDAWVKKGK. FDRI CTESDYEAFDASQDEFI MAFELELMKYLRLPS YI ESKVTEVLPKNLYI HSGKNI DDLAAWVTTSK. FNGVCTESDYEAFDASQDHFI LAFELEVMKFLGLPS YI EKKLMQALKPNFYI HSGKGLDELNEWVRTRG. FTGI CTESDYEAFDASQDHFI LAFELQI MKFLGLPE

DLI. DY. FIK. *LGSKLG*F.IMRF*GBASTFLFNT*ANMLFTF*RY. *. G. E. I. FAGDDMCA*#RL.. DLI EDYKFIKTSLGSKLGNFAIMRFSGEASTFLFNTLANMLFTFMRYNIRGDEFICFAGDDMCASRRLQP DLI ADYTFIKTHLGSKLGSFAIMRFTGEASTFLFNTMANMLFTFLRYDLNGREAICFAGDDMCANSRLKV DLI LDYEFIKIHLGSKLGSFSIMRFTGEASTFLFNTMANMLFTFLRYELTGSESIA<u>FAGDDMCA</u>NRRLRL

SPaV-1 Rep-II

日日日日 - KPTFCGWHLCPDGI YKKPQLVLERMCI AKEMNNLSNCI DNYAI - - PTFCGWGLCEHGVFKKPDLVLERLQI ARETRNLENCI DNYAI - - PTFCGWCLFKEGI FKKPQLI WERI CI AREMGNLENCI DNYAI ...FL..I.LKAKVQF#...---PTFCGW.L...G#*KKP#L..ER.#IA*E#.NL*NCIDNYAI I KKF AHF L DKL KL KAK VQF VQF VN· TNRFSNFLDKI KLKAKVÕFTÄT---KTEHEGFLNMI CLKAKVÕFVSN-ASPV Rep-11 PVM Rep-II

RSPaV-I Rep-II

OTSCIENT

VRNKH VS CAYKMGENLNLYLTPQEVDAHYNCVRF1 VQHNH VS YAYRLGELAI EMMTEEEVEAHYNCVRFLVRNKH V+#AY.#GE.#...#+#EV.A+YNCVR+.V###H V A Y A Y K L G E K A V N R M D E E E V A A F Y N C V R I I SSPaV-1 Rep-II

SPV Rep-II

PVM Rep-II

Consensus

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ASPV Rep-II VM Rep-II

RSPaV-1 Rep-II

ASPV Rep-II

PVM Rep-11

Consersus

V
K
_
76
6
H

M # L * . F L P. V. H. VPG*GK##L!### # . * # . A. T#GV*#. # * # . G. *! * MDV! VDLLYKYKFERLSNKL-VCP! VVHCVPGAGKSSL!RELLELDSRFCAYTAGVEDQPRLSGNW!RKW MDV!VDLLYKYKFERLSNKL-SDP!VVHAVPGSGKTTL!KQAL!RNN!EAVTFGVPEKAN!HGTY!KKA MNNLVKALSAFEFVGVFSVL-KFPVV!HSVPGSGKSSL!REL!SEDENF!AFTAGVPDSPNLTGRY!KPY	#. G G# *LDEY**	# * # * # * G*. * * * # * # H E. * . * * # * * G. TF. # VT # # # #
M # L * . F L.	#. G G# *LDEY** S - GQQP EGKF VVL DEYTLL - RQGQRGRGNYS I LDEYLSG - I SP GCAVP GKVNI LDEYLS V - (S - 1)	SEKADL VQVS DI YTKD-P SSKEDI VEKKNI FQLI EP SDGLDKVI VGSPFTLDV-
Consensus PVM 25K ASPV 25K RSPaV-1 24.4K	Majority PVM 25K ASPV 25K RSPaV-1 24.4K	Majority VM 25K ASPV 25K

CGP KQS	8/16
MPFQQ PANWAKTI TPLTVGLGI GLVLHFLRKSNLPYSGDNI HQFPHGGRYRDGTKSI TYCGPKQS	***S****

.... PPDYSKSVFPI AVGI AVAVVLFTLTRSTLPQVGDNI HNLPHGGNYQDGTKRI SYCGPRDS

ASPV 12K

PVM 12K

CONSTIN

TPPPDFTKVYLSAALGVSLALVVWLLIRSTLPVVGDRDHNLPHGGWYRDGTKSVFYNSP··

FPSSGIFGQSENFVPLMLVIGLIAFIHVLSVWN-SGLGRNCNCHPNPCSCRQQ

VM 12K 1SPV 12K

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FIG. 5C

I SQGQS...DCVVLI TGES VRVQGCRI DGEFGS...VLSKLKPFG AQQL YMS NS SQCTI VI TGES VS VVGCVYS EAFI EL VKGLKP YYHPLG IDS SSG. NHQGCFI RATGESI LI ENCGP S EALAS TVKE VLGGLKALG MI VYVLVGLSAFCI V···LY
MFPRSGLGLAVAAAVVAYLV
MYCLFGI·LVLVGI VI AI QI CORSEINS
PVM 7K
ASPV 8K
RSPaV-1 8.4K

Majority CGSFRS
PVM7K CGSFRS
ASPV 8K

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.. TLR#. C.. YA*. * WN. . L. * . . PPA* W#. * #F. #A*FD*F. * V. . . . # * * P. . G. . R. PT. * E 188) DAETL RRVCRL YAP VT WNHML THNAP PAE WAAMGF QYEDRFAP FDCFDYVENT AA VQP LEGLI RRP TP RE 301) EGCTLRQYCAFYAKHVWNLMLQTQSPPANWVGKEFKFETRYAAFDFFFGVESTASLEPADGLIRLPTQAI 142) EVTTLRRFCMYYAKI VWNI HLETGI PPANWAKKGFNENEKFAAFDFFLGVTDESALEPKGGI KRAPTI RSPaV-1 CP ASPV CP

RVANATSKEI QMYRI RSMEGTQAVNFGEVTGGKI GP--- KPVLSI - RK MVANI ASFEVÕVLRQAMAEGKRSSNLGEI SGGTAGALI NNPFSNVTHE KVAHNTHKDI AL. RGANRNQVFSSLNAEVTGGMNGPELTRDYVKSNRI VA*, *, *, *, *, . R. #., **, . ##*, *E. . GG. . G. . . . * VM CP

FIG.5D

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Consensus TTAGTTAATTAATTCTCCTGCA-TTCAAT RSPav-1 3UTR GGATGACGAAGTCAGCGACAATTCCGC Consensus CG. GG C. T GT. T CO GGCGGCTATGT GTGTTTT RSPav-1 3UTR CG GGCGGCTATGT TTT Consensus ASPV 3UTR ASPV 3UTR AAGAAGATTTGGTGTTTTTATAGTTTT RSPav-1 3UTR AAGAAGATTTGGTGTTTTTATAGTTTT	TTAGTTAATTAATTCTCCTGCA-TTCAATTTCAGTACTTATGCTTTTTAGTAAGTTGATCCCAACCTAAC GGATGACGAAGTCAGCGACAATTCCGCAGTCCAATAATTCCCCGATTTCAAGGCTGGGTTAAGCCTGTT GGATGACGAGTCAGCGACAATTCCGCAGTCCAATATTCTGT CG GGCCGCTATGTGT.TTT.CATGCTTAGCTTATTTTTGT CG GGCCGCTATGTTAG.TTTTC TTTTAG.TTTTC AAGAAGATTTGGTGTTTTTAAATTCATTCCATCCTAATCCTAATTAAT
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FIG. 6A

Consensus	
PVM 3' UTR RSPav-1 3'UTR	GGATGACGAAGTCAGCGACAATTCCGCAGTCCAATAATTCCCCGATTTCAAGGCTGGGTTAAGCCTGTTCGT
Consensus PVM 3' UTR RSPaV-1 3'UTR	GGATACCGTACTATAGGTATTCCCTTTCCTATTTAATATATAACGTGTG. A AAA. A GGAATACCGTACTAATACCTATTCCCTTCCATGCTAAAAGGTGTGGAAAGAAA
Consensus FVM 3'UTR ESPaV-1 3'UTR	A. A. TTGGT. T. T TAT TTTT

FIG. 6B

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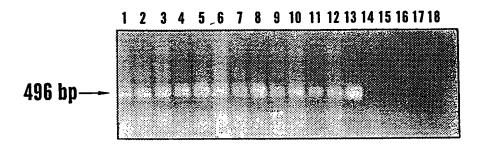


FIG. 7A

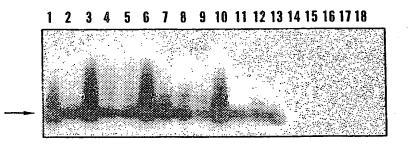


FIG. 7B

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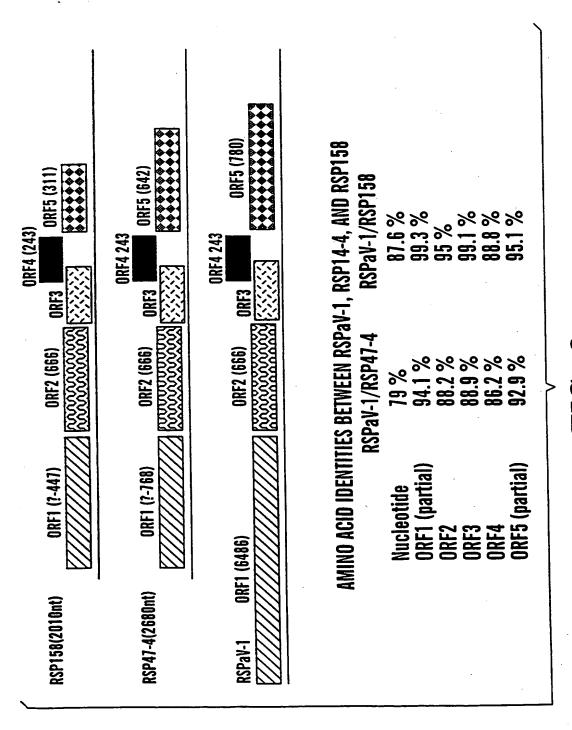
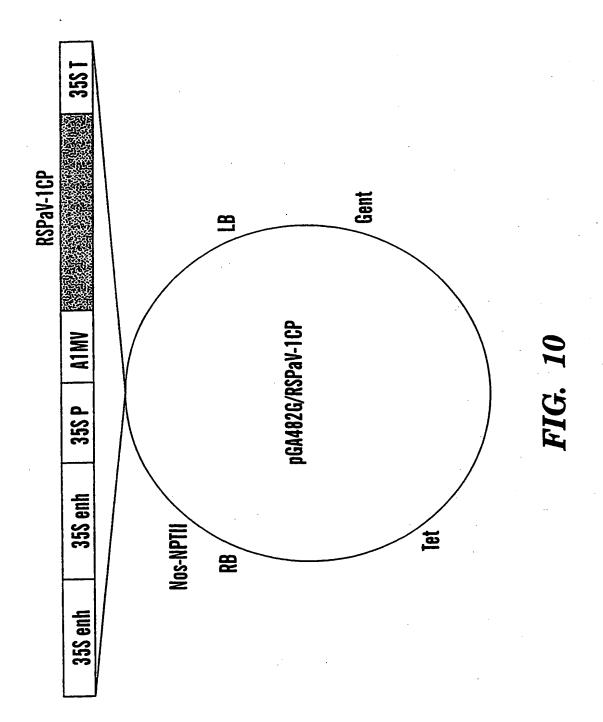


FIG. 8

FIG. 9

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.C. GGT. AA. G. TGG. CA. TGTGT. ATTTT. GA. GAGGT. CAGTTGTTTCC. CC. GGA. A. ATCGAT. T G. T CTT. T. A. ACG. AG. GA. GCTT	GCAGGAITTGAAGGCTGGGCACTGTGTGATTTTTGATGAGGTCCAGTTGTTTCCTCCTGGATACATCGATCTATGGTTGCTTATTATACGTAGTGATGCTT [ACAGGAITTGAAGACTGGTCATHTTGATGAGGGTCCAGTTGTTTCCTCCTGGATACATCGATCTGTTTTTGCTTATTATACGTAGTGATGCTT RC GCAGGTGTAAAAACTGGTCACTGGTGATTTTTGCAGGGTCCAGTTGTTTCCCCCTGGATACATCGATTTGCTTTTTTATACGTAGTGATGCTT GCAGGTATAAAAGTTGGTCACTGGTGAATTTTGCAAAAGAGCTCCAGTTGTTTCCCCCTGGATACATGTTTGCTTATAAACGTAGTGATGCTT 1RC GCAGGAATTGAAGGCTGGCCACTGTGTTTTGAAAAGGTCCAGTTGTTTCCTCCTGGATACATCGATCATTATTATACGTAGTGATGCTT GCAGGAATTGAAGGCTGGCCACTGTGTTTTTGATGAGGGTCCAGTTGTTTCCTCCTGGATACATCGATCTATGTTATACGTAGTGATGCTT GCAGGAATTGAAGGCTGGCCACTGTGATTTTTGATGAGGGTCCAGTTGTTTCCTCCTGGATACATCGATCTATGTTTATACGTAGTGATGCTT GCAGGAATTGAAGGCTGGCCACTGTGATTTTTGATGAGGGTCCAGTTGTTTCCTCCTGGATAGATCGATC	T.ATTT.ACT.GC.GGTGA.CCATG.CAGCACATATGA.TC.CA.AA.GATCG.GC.ATTTTGGG.GCTGAGCAGAGTGACATACT.A.A.T.CTTGA	1 ICATTICACTIGOGGGTGATCCATGTCABAGCACATATGATTCGCAAAAGGATCGGGCAATTTTGGGGCGCTGAGCAGGTGACATACTTAGAATGCTTGA 1 ICATTICACTTGCTGGTGATCCATGTCAGAGCACATATGATTGAGAGCATTTTGGGGCGCTGAGCAGGTGACATACTTGAGGTGCTTGA RC TCATTTCACTTGCTGGTGACCATGTGAGCACATATGAGTGACAAAAGATTTTGGGCGCTGAGCAGGTGACATACTTGAGATACTTGA TCATTTAACTTGCTGGTGAGCCATGTGAGCACATATGAGTGAAAAGATCGCGGGGATTTTGGGCGCTGAGCAGGTGACATATGAATACTTGA 1RC TCATTTCACTTGGGGGTGATCCATGTGAAAGATGCTGCAAAAGGATCGGGCAATTTTGGGCGCTGAGCAGGTGACATATAGAGTGCTTGA TCATTTCACTTGGGGGTGATCCATGTCAAAAGCATATGATTGGCAGCAAAAGGATTTTGGGCGCTGAGCAGAGTGACATAGAATGCTTGA TCATTTCACTTGCGGGTGATCCATGCAGAGCACATAGATTGGCAGAAAGGATCGGGCAATTTTGGGGGCTGAGCAGAGTGACATAGAATGCTTGA TGATTTCACTTGCGGGTGATCCATGCAGAGCACATAGATTGAGAAGGATCGGGCAATTTTGGGGCGCTGAGCAGAGTGACATACTTAGAGTTGCTTGA TGATTTCACTTGCGGGTGATCCATGGAGCACATAGATTCGCAAAAGGATCGGGCAATTTTGGGCGCTGAGCAGAGTGACATACTTAGAGTTGCTTGA TGATTTCACTTGCCGGTGATCCATGAGAGCACATAGAGTTCGCAAAAGGATCGGGCAATTTTGGGCGCTGAGCAGAGTGACATACTTAGAGTTGCTTGA	. GG. AA . AC. TATAGGTA . AACAT . GAAAGCAG G. TTTGTGAA . CCAATGTT . GAAT GACT . CCATGTCA . TTCAAAAA . GGG C. ATGACTGC	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	. GC. TT. GCTGATTATGC.AT.TT	CGCTTTGCTGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCGTTTTGGTTTCCAGTTTTGAGGAG ACCTTTGCTGATTATGCAATCTTCCACAATATGCATGACTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCTGTTTGGATTTTGAGGAG RC TTGCTTTGCTGATTATGCCATTTTCCACAATATGCATGATTTCCTTGGAGGTCAAAAGGTCCTTTGGATGCCATTTGGATTTTGAGGAG TGCTTTGCTGATTATGCCATTTTGCATATGCAAATATGCATGATTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCATTTTGATTTTGAGGAG CCTTTGCTGCTGATTATGCAATCTTCCATAATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCGTTTTGGTTTCCAGTTTTGAGGAG CGCTTTGCTGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCGTTTTGGTTTCCAGTTTTGAGGAG CGCTTTGCTGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCGTTTTGGTTTCCAGTTTTGAGGAG CGCTTTGCTGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCTTTGGATGCCGTTTTGGTTTTCAGTTTTGAGGAG CGCTTTGCTGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAGGTCCCTTTTGGATGCCGTTTTGGTTTTCAGTTTTGAGGAG CGCTTTGCCGATTATGCAATCTTCCATAATATGCATGACTTTCTCCTGGCGAGGTCAAAAAGGTCCCTTTTGGATGCCGTTTTTGGTTTTCCAGTTTTGAGGAG CGCTTTGCCTTTGCCGATTATGCAAATGCATAATATGCATGACTTTCCTTTGGATGCCGTTTTTGGATTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGTTTTGAGGAG
Consensus	140/94-1917+R1 140/94-2417+R1 140/94-213+F1RC 140/94-6417+R1RC 140/94-6417+R1RC 140/94-573+F RSPav-1	Consensus	140/94-19T7+R1 140/94-24T7+R1 140/94-2T3+F1RC 140/94-64T7+R1RC 140/94-6T3+F 140/94-6T3+R RSPaV-1	Consensus	140/94-1917+R1 140/94-2417+R1 140/94-213+F1RC 140/94-6477+R1RC 140/94-673+F 140/94-7217+R1 RSPaV-1	Consensus	140/94-1917+R1 140/94-2417+R1 140/94-213+F1RC 140/94-6417+R1RC 140/94-613+F 140/94-17+R1 RSPav-1

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. INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/10391

	SIFICATION OF SUBJECT MATTER			
	C07H 21/04; C07K 1/00; C12Q 1/68			
US CL :	435/6; 530/350; 536/24.3 International Patent Classification (IPC) or to both t	national classification and IPC		
F		indicated Classification and 17 C		
	DS SEARCHED	1 hardway Cardan markala		
Minimum do	ocumentation searched (classification system followed	by classification symbols)		
U.S. : 4	35/6; 530/350; 536/24.3			
Documentati	on searched other than minimum documentation to the	extent that such documents are included in the fields searched		
Electronic d	ata base consulted during the international search (na	ame of data base and, where practicable, search terms used)		
APS, AGF	RICOLA, BIOSIS, EMBASE, WPIDS			
search terr	ms: rupestris stem pitting			
	UMENTS CONSIDERED TO BE RELEVANT			
Category*	Citation of document, with indication, where ap	propriate, of the relevant passages Relevant to claim No.		
Y	SALATI et al. Detection of Grapev Leafroll, Corky Bark, and Rupestris ELISA and dsRNA Techniques. Ame Viticulture. 1994, Vol. 45, No. 3, pa	vine Viruses Associated with Stem Pitting Using F(ab') ₂ - crican Journal of Enology and		
Y	AZZAM et al. Detection of dsRi Symptoms of Rupestris Stem Pitting Encountered. Plant Disease. Septer pages 960-964, see especially pages 9	Disease and the Variabilities nber 1991, Vol. 75, No. 9,		
X Furth	ner documents are listed in the continuation of Box C	See patent family annex.		
• Sp	ecial estegories of cited documents:	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand		
.V. qo	cument defining the general state of the art which is not considered be of particular relevance	the principle or theory underlying the invention		
1	tier document published on or after the international filing data	"X" document of particular relevance; the claimed invention cannot be considered povel or cannot be considered to involve an inventive step		
·L· do	cument which may throw doubts on priority claim(s) or which is od to establish the publication date of another citation or other	when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be		
•0• do	ecial reason (as specified) rument referring to an oral disclosure, use, exhibition or other ans	considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art		
P do	ement publishes prior to the international filing date but later than priority date claimed	*&* document member of the same patent family		
	actual completion of the international search	Date of mailing of the international search report 1 4 OCT 1998		
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Authorized officer THANDA WAI				
_	L, D.C. 20231	Telephone No. (703) 308-0196		
i racsimile N	o. (703) 305-3230	1 010 pilotio 140.		

- INTERNATIONAL SEARCH REPORT

International application No.
PCT/US98/10391

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C (Continua	tion). DOCUMENTS CO	NSIDERED TO BE RELEVANT		
Category*	Citation of document,	with indication, where appropriate, of the relevant	passages	Relevant to claim No
Y		Double-Stranded RNA from Rupestris apevines. Vitis. 1989, Vol. 28, pages s 140-142.		51-63
Y		BECTON, DICKINSON & COMPAN pages 4-6 and 10-19.	TY). 01	51-63
A	Investigation of Gra	The Use of In Vitro Cultures in the apevine Virus-Like Diseases. Canadia 1990, Vol. 12, No. 3, page 337, see		1-5, 14-15, 51-63
A,P	Sources from Italy.	terization of Grapevine Rugose Wood Plant Disease. November 1997, Vol 2, see entire document.		1-5, 14-15, 51-63
A	Conidia of the Grap	tection of dsRNA from Cleistothecia a be Powdery Mildew Pathogen, Uncinutember 1991, Vol. 75, No. 9, pages 96 t.	la necator.	1-5, 14-15, 51-63
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_ INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/10391

Box I C	bservations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This inten	national report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
ı. 🗀	Claims Nos.:
ш	because they relate to subject matter not required to be searched by this Authority, namely:
2.	Claims Nos.:
L	because they relate to parts of the international application that do not comply with the prescribed requirements to such
	an extent that no meaningful international search can be carried out, specifically:
3.	Claims Nos.:
٣ ا	because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	mational Searching Authority found multiple inventions in this international application, as follows:
Pl	ease See Extra Sheet.
1. X	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite paymen of any additional fee.
	·
3.	As only some of the required additional search fees were timely paid by the applicant, this international search report cover only those claims for which fees were paid, specifically claims Nos.:
	only mose claims for which fees were paid, specifically claims Nos
4.	No required additional search fees were timely paid by the applicant. Consequently, this international search report i
لب	restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	on Protest The additional search fees were accompanied by the applicant's protest.
_ ,	No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

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BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be searched, the appropriate additional search fees must be paid.

Group I, claims 1-5 and 14-15, drawn to the replicase protein of Rupestris stem pitting associated virus, the first product.

Group II, claims 1-2, 6-8, and 14-15, drawn to the coat protein of Rupestris stem pitting associated virus, the second product.

Group III, claims 1-2 and 9-15, drawn to a triple gene block protein of Rupestris stem pitting associated virus, the third product.

Group IV, claims 16-17, drawn to an isolated RNA molecule encoding the replicase protein of Rupestris stem pitting associated virus, the fourth product.

Group V. claims 16-17, drawn to an isolated RNA molecule encoding the coat protein of Rupestris stem pitting associated virus, the fifth product.

Group VI, claims 16-17, drawn to an isolated RNA molecule encoding a triple gene block protein of Rupestris stem pitting associated virus, the sixth product.

Group VII, claims 18-19, 20-22, 33-40, and 62-63 drawn to an isolated DNA molecule encoding the replicase protein of Rupestris stem pitting associated virus, the seventh product.

Group VIII, claims 18-19, 23-25, 33-40, and 62-63 drawn to an isolated DNA molecule encoding the replicase protein of Rupestris stem pitting associated virus, the eighth product.

Group IX, claims 18-19, 26-40, and 62-63 drawn to an isolated DNA molecule encoding the replicase protein of Rupestris stem pitting associated virus, the ninth product.

Group X, claims 41-45, drawn to a method of imparting Rupestris stem pitting associated virus resistance to a Vitis scion or rootstock comprising transforming the Vitis scion or rootstock with a DNA molecule encoding replicase, the first method of using the seventh product.

Group XI, claims 41-45, drawn to a method of imparting Rupestris stem pitting associated virus resistance to a Vitis scion or rootstock comprising transforming the Vitis scion or rootstock with a DNA molecule encoding coat protein, the second method of using the eighth product.

Group XII, claims 41-45, drawn to a method of imparting Rupestris stem pitting associated virus resistance to a Vitis scion or rootstock comprising transforming the Vitis scion or rootstock with a DNA molecule encoding a triple block protein, the third method of using the ninth product.

Group XIII, claims 46-50, drawn to an antibody for the replicase protein of Rupestris stem pitting associated virus and to a method of using the antibody in an immunoassay, the tenth product and the fourth method of using the tenth product.

Group XIV, claims 46-50, drawn to an antibody for the coat protein of Rupestris stem pitting associated virus and to a method of using the antibody in an immunoassay, the eleventh product and the fifth method of using the eleventh product.

Group XV, claims 46-50, drawn to an antibody for a triple gene block protein of Rupestris stem pitting associated virus and to a method of using the antibody in an immunoassay, the twelfth product and the sixth method of using the twelfth product.

Group XVI, claims 51-54, drawn to a method for detecting Rupestris stem pitting associated virus in a sample by using a probe in a nucleic acid hybridization assay, the seventh method of using the products of Groups IV-IX.

Group XVII, claims 55-58, drawn to a method for detecting Rupestris stem pitting associated virus in a sample by using a gene amplification detection procedure, the eighth method of using the products of Groups IV-IX.

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Group XVIII, claims 59-61, drawn to oligonucleotide primers described by SEQ ID NO:41-54, the thirteenth product.

The inventions listed as Groups I-XVIII do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: the products of Groups I-IX, XII-XV, and XVIII are distinct. The methods of Groups X-XVII do not utilize the product of Group I. The search of Group I is limited to the replicase protein. Azzam et al. (1991) teach the detection of dsRNA in grapevines showing symptoms of Rupestris stem pitting disease. The reference teaches the RNA molecules of Groups IV-VI. Therefore, the special technical feature does not hold. PCT Rule 13 does not provide for multiple products or multiple methods of using within a single application (37 CFR 1.475(d)).

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